

FIG. 1a

Membranes from	RAW	264.7	P815	-
Affinity column	gp96	SA	gp96	.
212 _. ×				
116 =				Ş
83 ⊭			:	the same of the sa
51 ⊭				
35 ⊭				
28 ⊭				

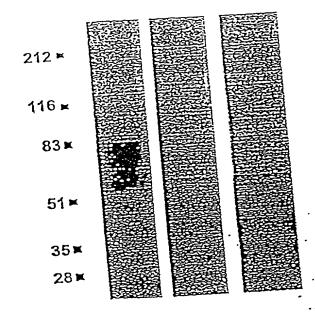
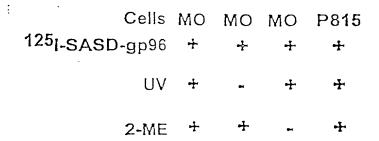


FIG. 1b



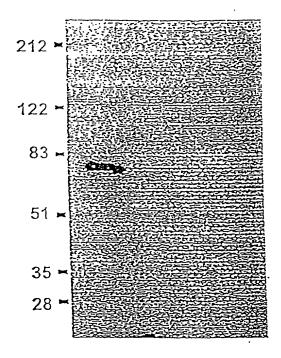


FIG. 1c

Pre-immune	Post-immune
PANJEA. Thacrophage	PANZEA. Tacrophage
1226	
83	
51 6	
35 cm	

FIG. 2a

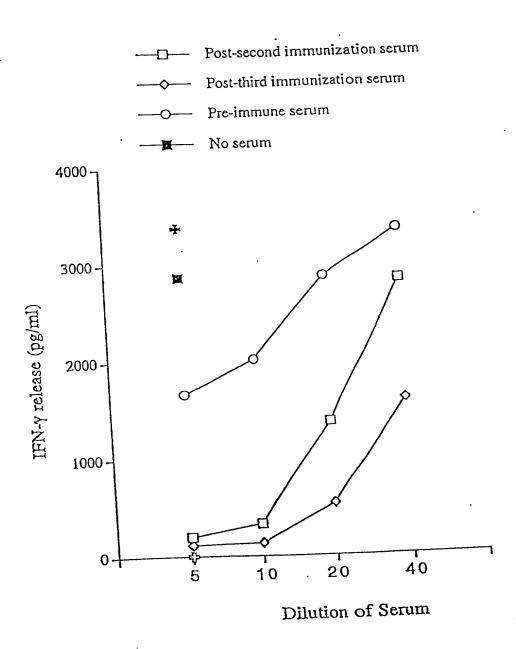


FIG. 2b

Sec	#	b	y	+1
G	1	58.1	-	10
G	2	115.1	1095.2	9
Α	3	186.2	1038.2	8
L	4	299.3	967.1	7
H	5	436.5	853.9	6
I	6	549.6	716.8	5
Y	7	712.8	603.6	4
Η	8	850.0	440.5	3
Q	9	978.1	303.3	2
R	10	-	175.2	1

FIG. 3a

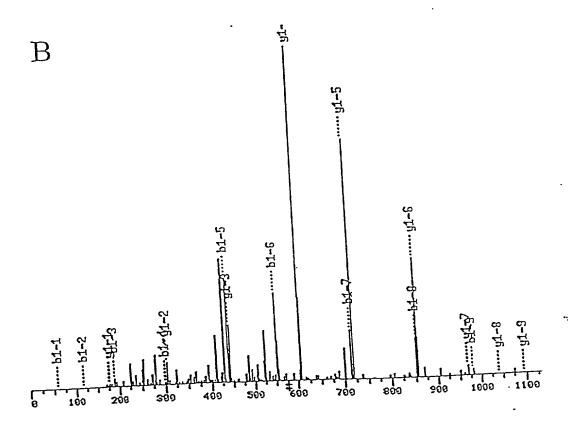
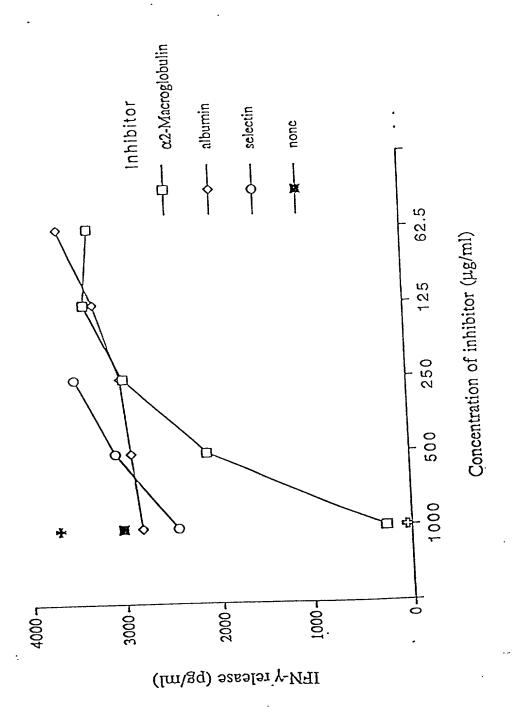


FIG. 3b

Position	MH+	Sequence
328-337	973.1753	SGFSLGSDGK (SCO 10 M:54) GIALDPAMGK (SCO 10 MO:55) GGALHIYHQR (SCO 10 MO:56) VFFTDYGQIPK (SCO 10 MO:57)



F1G. 2

Table 1. Specific binding of HSPs and α_2 -macroglobulin to primary cultures and cell lines of several histological origins*

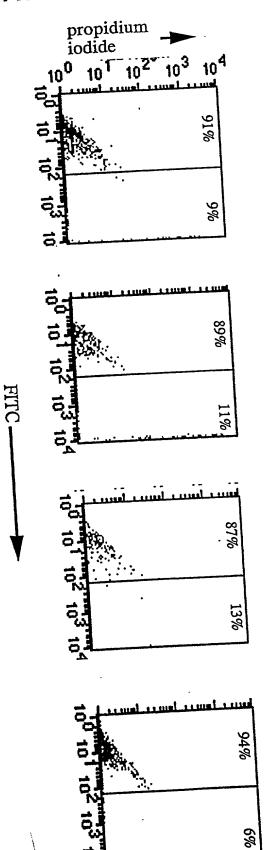
			**%	cells bir	nding wit	h FITC-lal	peled:
Cells	Celi type	Haplotype	α₂M	gp96	hsp70	hsp90	SA
B16	Melanoma	. b	0.1	3.5	6.4	8.0	0.3
CT26	Carcinoma	d	N/D	0.3	3.1	5.5	0.4
YAC-1	Lymphoma	b	0.1	3.1	23.0	5.0	0.2
EL4	T cell thymoma	b	0.1	2.9	3.0	6.6	1.0
Meth A	Sarcoma	d	0.1	0.1	1.5	0.9	0.5
PS-C3H	Fibrosarcoma	k	0.1	0.1	2.0	0.3	0.3
UV6139	Sarcoma	k	11	0.0	0.7	0.2	1.5
P815	Mastocytoma	d	0.1	1.1	1.7	0.7	0.2
Peritoneal cells	Macrophage	d	90	97	82	82	11
BM-DCs	Dendritic cells	b and d	+++#	+++	+++	+++	-
RAW264.7*	Macrophage	d	76	82	85	90	8.0
RAW309Cr.1*	Macrophage	bxd	0.1	0.1	0.1	0.1	0.1

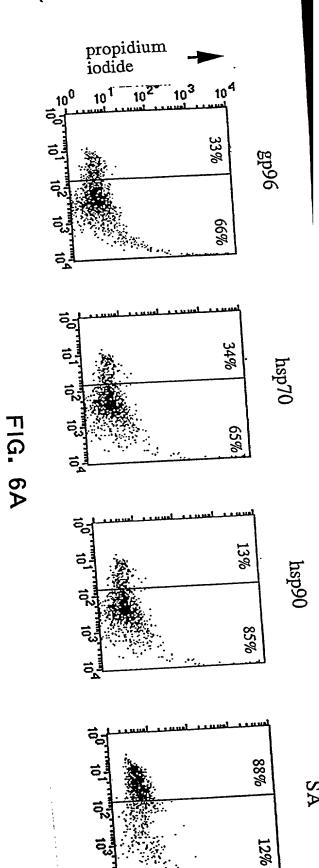
trend of the Will have de their W. H. Cont. W. H. Cont. West West Very Street S

the state that the state the state the

FIG.

6B





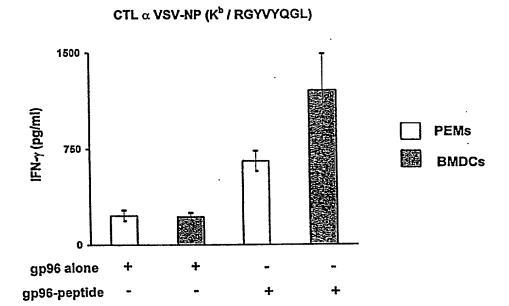


FIG. 7A

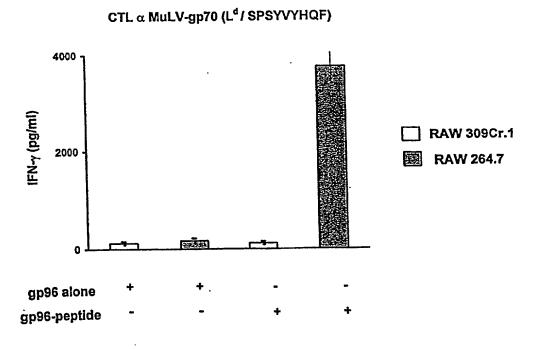
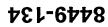
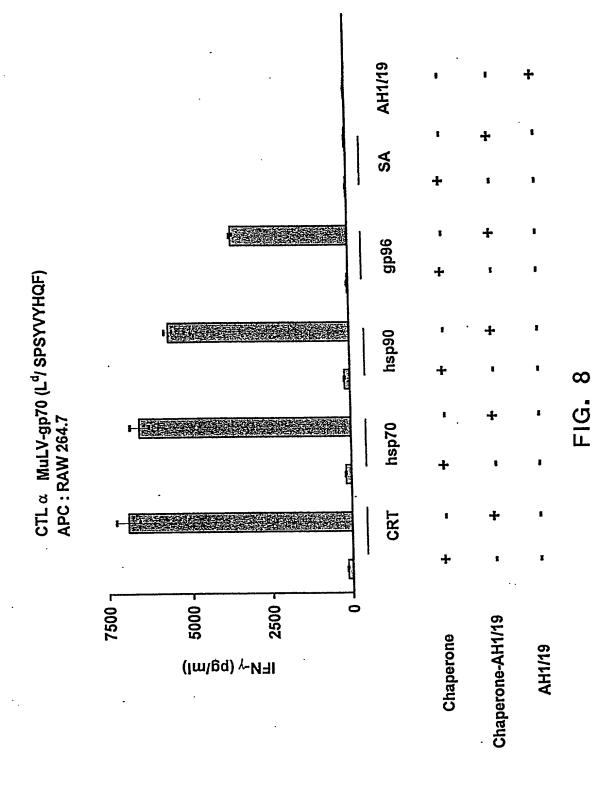


FIG. 7B

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APC: RAW 264.7 CTL against AH1 (Ld / SPSYVYHQF)

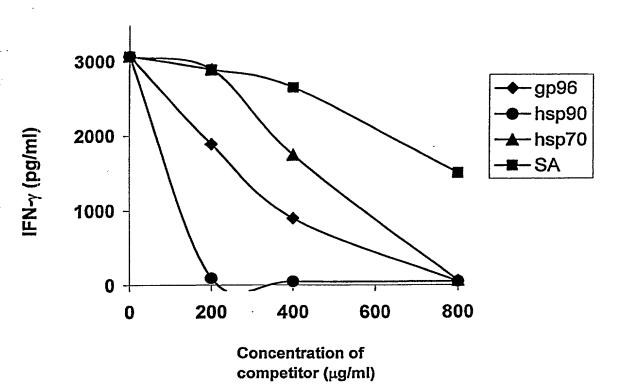


FIG. 9A



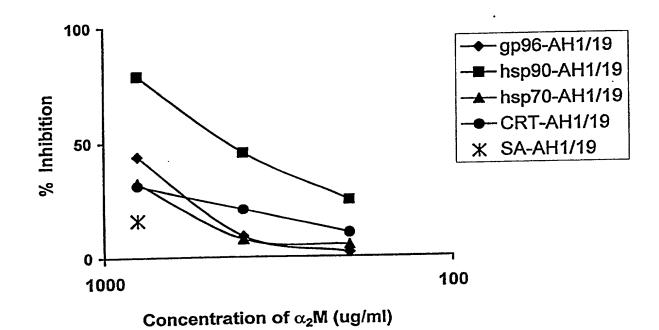
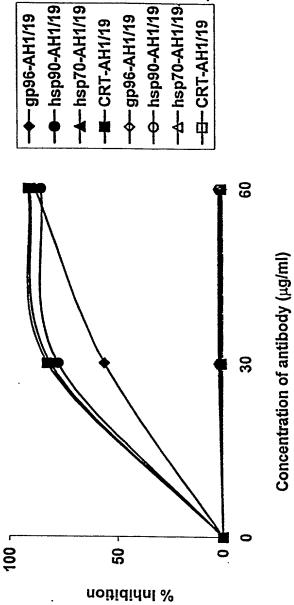


FIG. 9B



Isotype matched control Ab

Anti-CD91

FIG. 9C

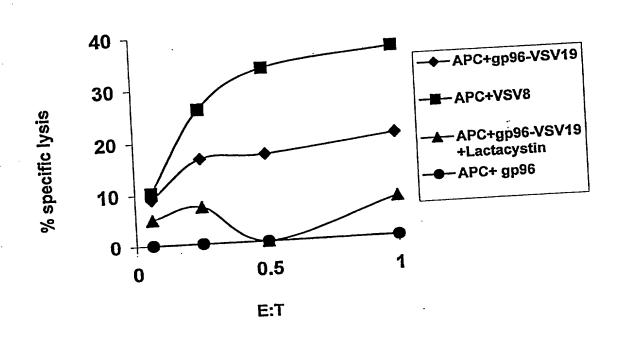


FIG. 10A

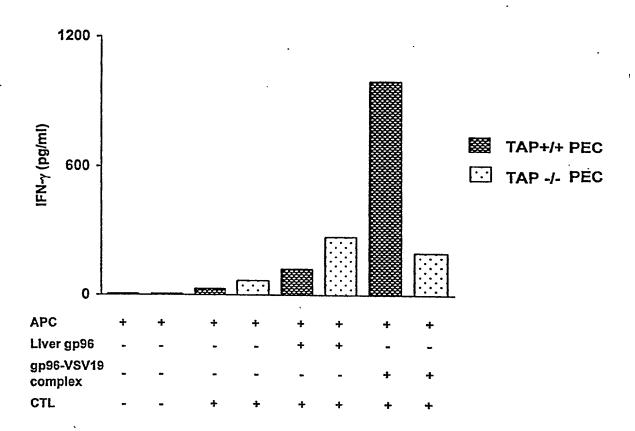


FIG. 10B

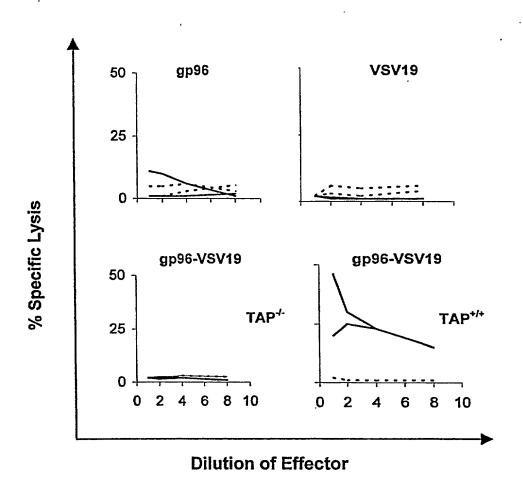


FIG. 10C

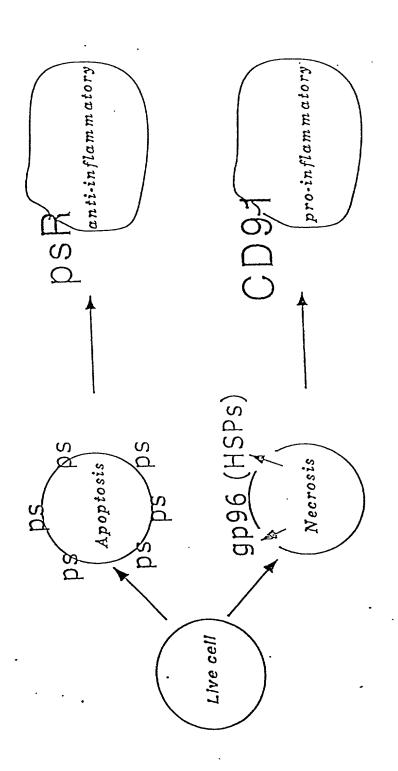


FIG. 11

GGCC CAAT GAGG CGCA CCTG GGGA	CCTA TGTG GGGA CCCG GTTC CCCC	CC A CA T GA G CG T GC T CC A	AGGC TTTT GAGC CAGC TTGC ATTG	ACCC GCAG GAGG AGGC TTAA GGGG	C CA C CG A GT C CT G GA G GG C C	TCGG GAGT AAAG TCCC AGGA CGAG ATG	GTCC CGGC CAGG AGGG TAAG GACA CTG	ACG TCC GGT GGC ATA AGA ACC	CCCC GAGA GAAG TCGG GAAG AGTA CCG	CCA TGG GGT AAC AGT ACA CCG	CCCC GGCT TCGA TGTA CGGG	CCACO GTGAO ATTTO CCATO GAGAO CAGAO CTG	CC C GC T GG G TT C GG A GG G	GCCT TCGC GGCA ACCT AGAT TGGG CTC		60 120 180 240 300 360 420 471
											ATG Met					519
											Gl'n CAA					567
											TGC Cys					615
											GCC Ala 70					663
CCA Pro 75	AAT Asn	GAG Glu	CAC His	AGT Ser	TGT Cys 80	CTG Leu	GGG Gly	ACT Thr	GAG Glu	CTA Leu 85	TGT Cys	GTC Val	CCC Pro	ATG Met	TCT Ser 90	711
											GGC Gly					759
GCT Ala	CAC His	TGC Cys	CGA Arg 110	GAG Glu	CTC Leu	CGA Arg	GCC Ala	AAC Asn 115	TGT Cys	TCT Ser	CGA Arg	ATG Met	GGT Gly 120	TGT Cys	CAA Gln	807
CAC His	CAT His	TGT Cys 125	GTA Val	CCT Pro	ACA Thr	CCC Pro	AGT Ser 130	GGG Gly	CCC Pro	ACG Thr	TGC Cys	TAC Tyr 135	TGT Cys	AAC Asn	AGC Ser	855
AGC Ser	TTC Phe 140	Gln	CTC Leu	GAG Glu	GCA Ala	GAT Asp 145	Gly	AAG Lys	ACG Thr	TGC Cys	AAA Lys 150	Asp	TTT Phe	GAC Asp	GAG Glu	903
	Ser					Cys					ACC Thr					951
					Cys					Lev	CTG				Asn	999
CGC Arg	TCC Ser	TGC Cys	AAG Lys 190	Ala	AAG Lys	AAT Asn	GAG Glu	CCP Pro	Val	GAD A	CGG Arç	CCG Pro	CCF Pro 200	Val	CTA Leu	1047

FIG. 12A

(Sheet 22 of 91)

CTG Leu	ATT Ile	GCC Ala 205	AAC Asn	TCT Ser	CAG Gln	AAC Asn	ATC Ile 210	CTA Leu	GCT Ala	ACG Thr	TAC Tyr	CTG Leu 215	AGT Ser	GGG Gly	GCC Ala	1095
CAA Gln	GTG Val 220	TCT Ser	ACC Thr	ATC Ile	ACA Thr	CCC Pro 225	ACC Thr	AGC Ser	ACC Thr	CGA Arg	CAA Gln 230	ACC Thr	ACG Thr	GCC Ala	ATG Met	1143
				GCC Ala		Glu										1191
AGT Ser	GCT Ala	GCC Ala	CAG Gln	ACA Thr 255	CAG Gln	CTC Leu	AAG Lys	TGT Cys	GCC Ala 260	CGG Arg	ATG Met	CCT Pro	GGC	CTG Leu 265	AAG Lys	1239
GGC	TTT Phe	GTG Val	GAT Asp 270	GAG Glu	His	ACC Thr	ATC Ile	AAC Asn 275	ATC Ile	TCC Ser	CTC Leu	AGC Ser	CTG Leu 280	CAC His	CAC His	1287
GTG Val	GAG Glu	CAG Gln 285	ATG Met	GCA Ala	ATC Ile	GAC Asp	TGG Trp 290	CTG Leu	ACG Thr	GGA Gly	AAC Asn	TTC Phe 295	TAC Tyr	TTT Phe	GTC Val	1335
				GAC Asp												1383
				CTG Leu												1431
				ATG Met 335												1479
				CGC Arg												1527
				ATC Ile											GTC Val	1575 ·
		Leu		TAC Tyr			Asp					Tyr				1623
	Asp			GGG Gly							Ile					1671
				TAC Tyr 415	Gly					Glu					Ala	1719
				Asn					Gln					. Ile	CGA Arg	1767

FIG. 12A

(Sheet 23 of 91)

GTG Val	AAC Asn	CGG Arg 445	TTC Phe	AAC Asn	AGT Ser	ACT Thr	GAG Glu 450	TAC Tyr	CAG Gln	GTC Val	GTC Val	ACC Thr 455	CGT Arg	GTG Val	GAC Asp	1815
AAG Lys	GGT Gly 460	GGT Gly	GCC Ala	CTG Leu	CAT His	ATC Ile 465	TAC Tyr	CAC His	CAG Gln	CGA Arg	CGC Arg 470	CAG Gln	CCC Pro	CGA Arg	GTG Val	1863
		CAC His														1911
		ATC Ile														1959
		TCT Ser														2007
		CAT His 525														2055
		GGC Gly														2103
		GAG Glu														2151
		TTC Phe														2199
		ATT		Gly												2247 •
		GTG Val 605	Glu					Asp								2295
TGG	ACT Thr 620	GAT Asp	GAT Asp	GGC Gly	CCC	AAG Lys 625	Lys	ACC	ATT	AGT Ser	GTG Val 630	Ala	AGG Arg	CTG Leu	GAG Glu	2343
AAA Lys 635	Ala	GCT Ala	CAG Gln	ACC Thr	CGG Arg 640	Lys	ACT Thr	CTA Leu	ATT Ile	GAG Glu 645	Gly	AAG Lys	ATG Met	ACA	CAC His 650	2391 -
CCC Pro	AGO Aro	G GCC J Ala	ATI	GTA Val 655	Val	GAT Asp	CCA Pro	CTC Lev	AAT Asr 660	Gly	TGG Trp	ATG Met	TAC Tyr	TGG Trp 665	Thr	2439
				. Asp					Arç					Glu	AGG Arg	2487

FIG. 12A

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GCT T	Trp	ATG Met 685	GAC Asp	GGC Gly	TCA Ser	CAC His	CGA Arg 690	GAT Asp	ATC Ile	TTT Phe	GTC Val	ACC Thr 695	TCC Ser	AAG Lys	ACA Thr	2535
GTG (2583
TAC Tyr 715																2631
GGC 2																267 _. 9
TTC (2727
AGC Ser																2775
ACT Thr																2823
ATG Met 795																2871
												CCC Pro			CGC . Arg	2919
CAG Gln	TGT Cys	GCC Ala	TGT Cys 830	Ala	GAG Glu	GAC Asp	CAG Gln	GTG Val 835	TTG Leu	GAC Asp	ACA Thr	GAT Asp	GGT Gly 840	GTC Val	ACC Thr	2967
TGC Cys	TTG Leu	GCG Ala 845	AAC Asn	CCA Pro	TCC Ser	TAC Tyr	GTG Val 850	Pro	CCA Pro	CCC Pro	CAG Gln	TGC Cys 855	Gln	CCG Pro	Gly GGC	3015
CAG Gln	TTT Phe 860	Ala	TGT Cys	GCC	AAC Asn	AAC Asn 865	Arg	TGC Cys	ATC Ile	CAG Gln	GAG Glu 870	Arg	TGG	AAG Lys	TGT Cys	3063
GAC Asp 875	GGA Gly	GAC Asp	AAC Asn	GAC Asp	TGT Cys 880	Leu	GAC Asp	AAC Asn	AGC Ser	GAT Asp 885	Glu	GCC Ala	CCA Pro	GCA Ala	Leu 890	3111
TGC Cys	CAT	CAA Gln	CAC His	ACC Thr 895	Cys	CCC Pro	TCC Ser	GAC Asp	CGA Arg 900	Phe	AAC Lys	TG1 Cys	GAG Glu	AAC Asr 200	AAC Asn	3159
CGG Arg	TGT Cys	ATC Ile	910	Asr	CGC Arg	TGC Trp	CTC Lev	C TG1 L Cys 915	Asp	GGC GLY	G GAS	raa 1 Rac o	GAT Asp 920	Cys	GGC Gly	3207

FIG. 12A

(Sheet 25 of 91)

AAC Asn	AGC Ser	GAG Glu 925	GAC Asp	GAA Glu	TCC Ser	AAT Asn	GCC Ala 930	ACG Thr	TGC Cys	TCA Ser	GCC Ala	CGC Arg 935	ACC Thr	TGT Cys	CCA Pro	3255
												CCT Pro				3303
												GAT Asp				3351
												TTT Phe				3399
												AAC Asn				3447
	Gly					Glu					His	TCC Ser 1015				3495
Thr					Asn					Ile		GAG Glu				3543
TGT Cys 1035	GAT Asp	GGG Gly	GAC Asp	Asn	GAT Asp 1040	TGT Cys	GGG Gly	GAC Asp	Tyr	AGC Ser 1045	GAC Asp	GAG Glu	ACA Thr	His	GCC Ala 1050	3591
			Asn					Pro				TGC Cys	His		Asp	3639
GAG Glu	TTC Phe	Gln	TGC Cys 1070	Pro	CTA Leu	GAT Asp	Gly	CTG Leu 1075	Cys	ATC Ile	CCC Pro	Leu	AGG Arg 1080	Trp	CGC Arg	3687 ·
TGC Cys	Asp	GGG Gly 1085	Asp	ACC Thr	GAC Asp	Cys	ATG Met 1090	Asp	TCC Ser	AGC Ser	GAT Asp	GAG Glu 1095	Lys	AGC Ser	Cys	3735
Glu	GGC Gly 1100	Val	ACC Thr	CAT His	GTT Val	TGT Cys 1105	Asp	CCG Pro	AAT Asn	GTC Val	Lys	Phe	GLy	TGC Cys	AAG Lys	3783
	Ser					Ser					. Cys				AGC Ser 1130	3831
GAC Asp	TGT Cys	GAI Glu	A GAT	AAC Asr 1135	Ser	GAC Asp	GAC Glu	GAC i Glu	AA(1 Asi 114(ı Cys	GAC Glu	G GCC	CTC	G GC0 1 Al: 114!	C TGC a Cys 5	3879
AGG Arg	CCP Pro	CCO Pro	TC0 Ser 1150	r His	CCC Fro	TGC Cys	GCG Ala	C AAG A Asi 1159	n Ası	C ACC	C TC	r GT(C TG(L Cy: 116	s Le	G CCT u Pro	3927

FIG. 12A

(Sheet 26 of 91)

CCT Pro	Asp	AAG Lys 165	CTG Leu	TGC Cys	GAC Asp	Gly	AAG Lys 170	GAT Asp	GAC Asp	TGT Cys	GGA Gly 1	GAC Asp 175	GGC Gly	TCG Ser	GAT Asp	3975
Glu	GGC Gly 180	GAG Glu	CTC Leu	TGT Cys	Asp	CAG Gln 185	TGT Cys	TCT Ser	CTG Leu	Asn	AAT Asn 190	GGT Gly	GGC Gly	TGT Cys	AGT Ser	4023
CAC His 1195	AAC Asn	TGC C ys	TCA Ser	Val	GCC Ala 200	Pro	GGT Gly	GAA Glu	Gly	ATC Ile 205	GTG Val	TGC Cys	TCT Ser	Cys	CCT Pro 210	4071
CTG Leu	GGC Gly	ATG Met	Glu	CTG Leu 215	GGC Gly	TCT Ser	GAC Asp	Asn	CAC His 220	ACC Thr	TGC Cys	CAG Gln	Ile	CAG Gln 1225	AGC Ser	4119
		Ala					Cys				TGT Cys	Asp				4167
	Ser					Cys					GTC Val					4215
Gly					Ser					Lys	CTG Leu 1270					4263
				Glu					Asp		CAC His			Asp		4311
			Val					Asn			GCC Ala		Asp			4359
		Gln					Trp				GTA Val	Glu				4407
	Arg		Lys			Asp		Gly			ACC		Phe			4455
		Gln			Leu		Thr			Gly	CTG Leu 1350	Ala				4503
	Ala			Ile		Trp			Ser		CTG Leu			Ile		4551
			_		Gly					Thr	CTG				Asp	4599
				Arg					Asp		CGG Arg			/ Ile	CTG Leu	4647

FIG. 12A

TTT TGG ACA GAC TGG GAT GCC AGC CTG CCA CGA ATC GAG GCT GCA TCC Phe Trp Thr Asp Trp Asp Ala Ser Leu Pro Arg Ile Glu Ala Ala Ser 1405 1410 1415	4695
ATG AGT GGA GCT GGC CGC CGA ACC ATC CAC CGG GAG ACA GGC TCT GGG Met Ser Gly Ala Gly Arg Arg Thr Ile His Arg Glu Thr Gly Ser Gly 1420 1425 1430	4743
GGC TGC GCC AAT GGG CTC ACC GTG GAT TAC CTG GAG AAG CGC ATC CTC Gly Cys Ala Asn Gly Leu Thr Val Asp Tyr Leu Glu Lys Arg Ile Leu 1435	4791
TGG ATT GAT GCT AGG TCA GAT GCC ATC TAT TCA GCC CGG TAT GAC GGC Trp Ile Asp Ala Arg Ser Asp Ala Ile Tyr Ser Ala Arg Tyr Asp Gly 1455 1460 . 1465	4839
TCC GGC CAC ATG GAG GTG CTT CGG GGA CAC GAG TTC CTG TCA CAC CCA Ser Gly His Met Glu Val Leu Arg Gly His Glu Phe Leu Ser His Pro 1470 1475 1480	4887
TTT GCC GTG ACA CTG TAC GGT GGG GAG GTG TAC TGG ACC GAC TGG CGA Phe Ala Val Thr Leu Tyr Gly Gly Glu Val Tyr Trp Thr Asp Trp Arg 1485 1490 1495	4935
ACA AAT ACA CTG GCT AAG GCC AAC AAG TGG ACT GGC CAC AAC GTC ACC Thr Asn Thr Leu Ala Lys Ala Asn Lys Trp Thr Gly His Asn Val Thr 1500 1505 1510	4983
GTG GTA CAG AGG ACC AAC ACC CAG CCC TTC GAC CTG CAG GTG TAT CAC Val Val Gln Arg Thr Asn Thr Gln Pro Phe Asp Leu Gln Val Tyr His 1515 1520 1525 1530	5031
CCT TCC CGG CAG CCC ATG GCT CCA AAC CCA TGT GAG GCC AAT GGC GGC Pro Ser Arg Gln Pro Met Ala Pro Asn Pro Cys Glu Ala Asn Gly Gly 1535 1540 1545	5079
CGG GGC CCC TGT TCC CAT CTG TGC CTC ATC AAC TAC AAC CGG ACC GTC Arg Gly Pro Cys Ser His Leu Cys Leu Ile Asn Tyr Asn Arg Thr Val 1550 1555 1560	5127 ·
TCC TGG GCC TGT CCC CAC CTC ATG AAG CTG CAC AAG GAC AAC ACC ACC Ser Trp Ala Cys Pro His Leu Met Lys Leu His Lys Asp Asn Thr Thr 1565 1570 1575	5175
TGC TAT GAG TTT AAG AAG TTC CTG CTG TAC GCA CGT CAG ATG GAG ATC Cys Tyr Glu Phe Lys Lys Phe Leu Leu Tyr Ala Arg Gln Met Glu Ile 1580 1585 1590	5223
CGG GGC GTG GAC CTG GAT GCC CCG TAC TAC AAT TAT ATC ATC TCC TTC Arg Gly Val Asp Leu Asp Ala Pro Tyr Tyr Asn Tyr Ile Ile Ser Phe 1595 1600 1605 .	5271 -
ACG GTG CCT GAT ATC GAC AAT GTC ACG GTG CTG GAC TAT GAT GCC CGA Thr Val Pro Asp Ile Asp Asn Val Thr Val Leu Asp Tyr Asp Ala Arg 1615 1620 1625	.5319
GAG CAG CGA GTT TAC TGG TCT GAT GTG CGG ACT CAA GCC ATC AAA AGG Glu Gln Arg Val Tyr Trp Ser Asp Val Arg Thr Gln Ala Ile Lys Arg 1630 1635 1640	5367

FIG. 12A

(Sheet 28 of 91)

GCA TTT ATC AAC (Ala Phe Ile Asn (1645)	GGC ACT GGC GTG Gly Thr Gly Val 1650	Glu Thr Val Val	TCT GCA GAC TTG Ser Ala Asp Leu 1655	5415
CCC AAC GCC CAC (Pro Asn Ala His (1660	GGG CTG GCT GTG Gly Leu Ala Val 1665	G GAC TGG GTC TCC Asp Trp Val Ser 1670	CGA AAT CTG TTT Arg Asn Leu Phe	5463
TGG ACA AGT TAC O Trp Thr Ser Tyr 2 1675	Asp Thr Asn Lys 1680	Lys Gln Ile Asn 1685	Val Ala Arg Leu 1690	5511
_	Lys Asn Ala Val 695	Val Gln Gly Leu 1700	Glu Gln Pro His 1705	5559.
GGC CTG GTC GTC Gly Leu Val Val 1710	His Pro Leu Arc	Gly Lys Leu Tyr 1715	Trp Thr Asp Gly 1720	5607
GAC AAC ATC AGC A Asp Asn Ile Ser 1 1725	Met Ala Asn Met 1730	Asp Gly Ser Asn	His Thr Leu Leu 1735	5655
TTC AGT GGC CAG Phe Ser Gly Gln 1 1740	Lys Gly Pro Va] 1745	. Gly Leu Ala Ile 1750	Asp Phe Pro Glu	5703
AGC AAA CTC TAC Ser Lys Leu Tyr 1755	Trp Ile Ser Ser 1760	Gly Asn His Thr 1765	Ile Asn Arg Cys 1770	5751
	Ser Glu Leu Glu 775	val Ile Asp Thr 1780	Met Arg Ser Gln 1785	5799
CTG GGC AAG GCC . Leu Gly Lys Ala 1790	Thr Ala Leu Ala	a Ile Met Gly Asp 1795	Lys Leu Trp Trp 1800	5847
GCA GAT CAG GTG Ala Asp Gln Val 1805	Ser Glu Lys Met 1810	: Gly Thr Cys Asn)	Lys Ala Asp Gly 1815	5895
TCT GGG TCC GTG Ser Gly Ser Val 1820	Val Leu Arg Ası 1825	n Ser Thr Thr Leu 1830	Val Met His Met	5943
AAG GTG TAT GAC Lys Val Tyr Asp 1835	Glu Ser Ile Gli 1840	n Leu Glu His Glu 1845	Gly Thr Asn Pro 1850	5991
TGC AGT GTC AAC Cys Ser Val Asn 1	AAC GGA GAC TG Asn Gly Asp Cy: 855	T TCC CAG CTC TGC s Ser Gln Leu Cys 1860	CTG CCA ACA TCA Leu Pro Thr Ser 1865	6039
GAG ACG ACT CGC Glu Thr Thr Arg 1870	TCC TGT ATG TG Ser Cys Met Cy	T ACA GCC GGT TAC s Thr Ala Gly Tyr 1875	AGC CTC CGG AGC Ser Leu Arg Ser 1880	6087

FIG. 12A

GGA CAG Gly Gln					Gly '					Leu					6135
CAT GAG His Glu 1900				Gly					Pro						6183
GCC CTG Ala Leu 1915			Val					Leu					Asp		6231
CAT GCC His Ala		Asn					Trp					Leu			6279
ATC AGC Ile Ser	Arg					Gln					Asp				6327
AAC GGT Asn Gly					Glu					Asp					6375
AAC ATA Asn Ile 1980	Tyr	TGG Trp	ACG Thr	Asp	CAG Gln 1985	GGC Gly	TTC Phe	GAT Asp	Val	ATC Ile 1990	GAG Glu	GTT Val	GCC Ala	CGG Arg	6423
CTC AAT Leu Asn 1995	GGC Gly	TCT Ser	Phe	CGT Arg 2000	TAT Tyr	GTG Val	GTC Val	Ile	TCC Ser 2005	CAG Gln	GGT Gly	CTG Leu	Asp	AAG Lys 2010	6471
CCT CGG Pro Arc	GCC Ala	Ile	ACT Thr 2015	GTC Val	CAC His	CCA Pro	Glu	AAG Lys 2020	G) y	TAC Tyr	TTG Leu	Phe	TGG Trp 2025	ACC Thr	6519
GAG TGC Glu Tr	G GGT	CAT His 2030	Tyr	CCA Pro	CGT Arg	Ile	GAG Glu 2035	Arg	TCT Ser	CGC Arg	Leu	GAT Asp 2040	Gly	ACA Thr	6567 .
GAG AGA	A GTG g Val 2045	. Val	TTG Leu	GTT Val	Asn	GTC Val 2050	Ser	ATC	AGC Ser	TGG	CCC Pro 2055	Asn	GGC Gly	ATC Ile	6615
TCA GTA Ser Val 206	l Asp	TAT Tyr	CAG	Gly	GGC Gly 2065	Lys	CTC Leu	TAC Tyr	TGG	TGT Cys 2070	Asp	GCT Ala	CGG Arç	ATG Met	6663
GAC AA Asp Ly 2075	G ATO	C GAG ∋ Glu	G CGC	: ATC Ile 2080	Asp	CTG Lev	GAA Glu	Thr	GGC Gly 2085	/ Glu	AA G 12A 1	CGC Arg	G GAC G Glu	GTG Val 2090	6711
GTC CT Val Le	G TC	C AGO	AA1 Asr 2095	Asn	: ATG Met	GAT Asp	ATO Met	TTC Phe 2100	Se	C GTC	TC(C GTO	3 TT: 1 Phe 210:	e Glu	6759
GAC TT Asp Ph	C ATO	C TAC e Ty: 2110	r Trp	G AGT	GAC Asp	AGA Arg	A ACT g Th: 211	r His	C GCG S Ala	C AA' a Ası	r GGG	TC y Se: 212	r Il	C AAG e Lys	6807

FIG. 12A

(Sheet 30 of 91)

CGC Arg	Gly	TGC Cys 2125	AAA Lys	GAC Asp	AAT Asn	Ala	ACA Thr 130	GAC Asp	TCC Ser	GTG Val	Pro	CTG Leu 2135	AGG Arg	ACA Thr	GGC Gly	6855
Ile	GGT Gly 2140	GTT Val	CAG Gln	CTT Leu	Lys	GAC Asp 145	ATC Ile	AAG Lys	GTC Val	Phe	AAC Asn 2150	AGG Arg	GAC Asp	AGG Arg	CAG Gln	6903
AAG Lys 2155	GGT Gly	ACC Thr	AAT Asn	Val	TGC Cys 2160.	Ala	GTA Val	GCC Ala	Asn	GGC Gly 2165	GGG Gly	TGC Cys	CAG Gln	GIn	CTC Leu 2170	6951
TGC Cys	TTG Leu	TAT Tyr	CGG Arg	GGT Gly 2175	GGC Gly	GGA Gly	CAG Gln	Arg	GCC Ala 2180	TGT Cys	GCC Ala	TGT Cys	Ala	CAC His 2185	Gly GGG	6999
ATG Met	CTG Leu	Ala	GAA Glu 2190	Asp	GGG Gly	GCC Ala	Ser	TGC Cys 2195	CGA Arg	GAG Glu	TAC Tyr	Ala	GGC Gly 2200	TAC Tyr	CTG Leu	7047
CTC Leu	Tyr	TCA Ser 2205	GAG Glu	CGG Arg	ACC Thr	Ile	CTC Leu 2210	Lys	AGC Ser	ATC Ile	CAC	CTG Leu 2215	Ser	GAT Asp	GAG Glu	7095
CGT Arg	AAC Asn 2220	Lev	AAC Asn	GCA Ala	Pro	GTG Val 2225	CAG Gln	CCC Pro	TTT Phe	GAA Glu	GAC Asp 2230	Pro	GAG Glu	CAC His	ATG Met _.	7143
AAA Lys 2235	: Asr	GTC Nal	C ATC	GCC Ala	CTG Leu 2240	Ala	TTT Phe	GAC Asp	TAC Tyr	CGA Arc 2245	, Ala	A GGC	ACC Thr	Ser	CCG Pro 2250	7191
GG(ACC Thi	C CC	OAA 1	CGC Arg 2255	, Ile	TTC Phe	TTC Phe	AG1 Sei	GAC Asp 2260	o Ile	C CAG Hi:	C TTI s Phe	GGC Gly	AAC Asn 2265	ATC Ile	7239
CA(Gl:	G CAG	G AT	C AA e Asi 2270	n Asp	GAT Asp	GGC Gly	TCC Sei	G GGG Gl: 227	y Ar	G ACC	C AC	C ATO	C GT(e Va; 228	r er	A AAT a Asn	7287
GT Va	G GG 1 Gl	C TC y Se 228	r Va	G GAJ l Gli	A GGC u Gly	CTC Leu	G GC0 1 Ala 229	а Ту	T CA	C CG	T GG g Gl	C TGG y Tr 229	p As	C ACA	A CTG	7335
TA Ty	C TG	p Th	A AG	C TAG	C ACC	C AC c Thi 230	r Se	C AC r Th	C AT r Il	C AC e Th	C CG r Ar 231	d ur	C AC s Th	C GT r Va	G GAC l Asp	7383
CA G1 231	n Th	T CG	GC CC	A GG o Gl	G GC y Al 232	a Ph	C GA e Gl	G AG u Ar	G GA	G AC u Th 232	ır Va	C AT	C AC	C AT	G TCC t Ser 2330	7431 -
GI GI	SA GA Ly As	AC GA	AC CA sp Hi	C CC s Pr 233	o Ar	A GC g Al	C TI a Ph	T GT ne Va	CG CT al Le 23	eu As	AT GI	AG TG Lu Cy	GC CA	AG AA In As 234	C CTG in Leu 15	7,479
A: Me	rG TT et Pl	TC TC	GG AC rp Th	ir As	AT TG	G AA	C G!	AG Ci lu Lo 23:	eu H	AT CO	CA A	GC A1 er I1	rc A' le Mo 23	EC A	GG GCA	7527

FIG. 12A

(Sheet 31 of 91)

GCC CTA TCC GGA GCC AAC GTC CTG ACC CTC ATT GAG AAG GAC ATC CGC Ala Leu Ser Gly Ala Asn Val Leu Thr Leu Ile Glu Lys Asp Ile Arg 2365 2370 2375	7575
ACG CCC AAT GGG TTG GCC ATC GAC CAC CGG GCG GAG AAG CTG TAC TTC Thr Pro Asn Gly Leu Ala Ile Asp His Arg Ala Glu Lys Leu Tyr Phe 2380 2385 2390	7623
TCG GAT GCC ACC TTG GAC AAG ATC GAG CGC TGC GAG TAC GAC GGC TCC Ser Asp Ala Thr Leu Asp Lys Ile Glu Arg Cys Glu Tyr Asp Gly Ser 2400 2405 2410	7671
CAC CGC TAT GTG ATC CTA AAG TCG GAG CCC GTC CAC CCC TTT GGG TTG His Arg Tyr Val Ile Leu Lys Ser Glu Pro Val His Pro Phe Gly Leu 2415 2420 2425	7719
GCG GTG TAC GGA GAG CAC ATT TTC TGG ACT GAC TGG GTG CGG CGG GCT Ala Val Tyr Gly Glu His Ile Phe Trp Thr Asp Trp Val Arg Arg Ala 2430 2435 2440	7767
GTG CAG CGA GCC AAC AAG TAT GTG GGC AGC GAC ATG AAG CTG CTT CGG Val Gln Arg Ala Asn Lys Tyr Val Gly Ser Asp Met Lys Leu Leu Arg 2445 2450 2455	7815
GTG GAC ATT CCC CAG CAA CCC ATG GGC ATC ATC GCC GTG GCC AAT GAC Val Asp Ile Pro Gln Gln Pro Met Gly Ile Ile Ala Val Ala Asn Asp 2460 2465 2470	7863
ACC AAC AGC TGT GAA CTC TCC CCC TGC CGT ATC AAC AAT GGA GGC TGC Thr Asn Ser Cys Glu Leu Ser Pro Cys Arg Ile Asn Asn Gly Gly Cys 2475 2480 2485 2490	
CAG GAT CTG TGT CTG CTC ACC CAC CAA GGC CAC GTC AAC TGT TCC TGT Gln Asp Leu Cys Leu Leu Thr His Gln Gly His Val Asn Cys Ser Cys 2495 2500 2505	7959
CGA GGG GGC CGG ATC CTC CAG GAG GAC TTC ACC TGC CGG GCT GTG AAC Arg Gly Gly Arg Ile Leu Gln Glu Asp Phe Thr Cys Arg Ala Val Asn 2510 2520	8007
TCC TCT TGT CGG GCA CAA GAT GAG TTT GAG TGT GCC AAT GGG GAA TGT Ser Ser Cys Arg Ala Gln Asp Glu Phe Glu Cys Ala Asn Gly Glu Cys 2525 2530 2535	8055
ATC AGC TTC AGC CTC ACC TGT GAT GGC GTC TCC CAC TGC AAG GAC AAG Ile Ser Phe Ser Leu Thr Cys Asp Gly Val Ser His Cys Lys Asp Lys 2540 2550	8103
TCC GAT GAG AAG CCC TCC TAC TGC AAC TCA CGC CGC TGC AAG AAG ACT Ser Asp Glu Lys Pro Ser Tyr Cys Asn Ser Arg Arg Cys Lys Lys Thr 2555 2560 2565 2570	
TTC CGC CAG TGT AAC AAT GGC CGC TGT GTA TCC AAC ATG CTG TGG TGC Phe Arg Gln Cys Asn Asn Gly Arg Cys Val Ser Asn Met Leu Trp Cys 2575 2580 2585	8199 5
AAT GGG GTG GAT TAC TGT GGG GAT GGC TCT GAT GAG ATA CCT TGC AAC Asn Gly Val Asp Tyr Cys Gly Asp Gly Ser Asp Glu Ile Pro Cys Asi 2590 2595 2600	

						•										•
	ACT Thr 2					Gly					Arg					8295
Ile	GGG Gly 2620				Arg					Val						8343
	GAT Asp			Asn					Asp					Phe		8391
	GGC Gly		Lys					Gln					Thr			8439
	TAC Tyr	Ala					Cys					Asp				8487
TAC Tyr	AGC Ser	GAT Asp 2685	Glu	CGT Arg	GAC Asp	Cys	CCA Pro 2690	GGT Gly	GTG Val	AAG Lys	Arg	CCT Pro 2695	AGG Arg	TGC Cys	CCG Pro	8535
Leu	AAT Asn 2700	TAC Tyr	TTT Phe	GCC Ala	Cys	CCC Pro 2705	AGC Ser	GGG Gly	CGC Arg	Cys	ATC Ile 2710	CCC Pro	ATG Met	AGC Ser	TGG Trp	8583
ACG Thr 2715	TGT Cys	GAC Asp	AAG Lys	Glu	GAT Asp 2720	GAC Asp	TGT Cys	GAG Glu	Asn	GGC Gly 2725	GAG Glu	GAT Asp	GAG Glu	Thr	CAC His 2730	8631
TGC Cys	: AAC : Asn	AAG Lys	Phe	TGC Cys 2735	Ser	GAG Glu	GCA Ala	Gln	TTC Phe 2740	GAG Glu	TGC Cys	CAG Gln	Asn	CAC His 2745	CGG Arg	8679
TG1 Cys	T ATC	Ser	AAG Lys 2750	Gln	TGG Trp	CTG Leu	Cys	GAC Asp 2755	Gly	AGC Ser	GAT Asp	GAT Asp	TGC Cys 2760	Gly	GAT Asp	8727 .
GG(Gly	TCC Ser	GAT Asp 2765	Glu	GCA Ala	GCT Ala	His	TGT Cys 2770	Glu	GGC Gly	AAG Lys	ACA Thr	TGT Cys 2775	Gly	CCC Pro	TCC Ser	8775
TC0 Sea	TTC Phe 2780	Sei	TG1 Cys	CCC Pro	GGC Gly	ACC Thr 2785	His	GTG Val	TGT Cys	GTC Val	CC1 Pro 2790	Glu	G CGC	TGG Trp	CTC Leu	8823
TG: Cy: 279:	s Asp	GGG Gly	C GAG y Ası	C AAC D Lys	G GAC S Asp 2800	Cys	ACC Thr	C GAT	GGC Gly	GCG Ala 2809	. Asp	GAC	G AGT	GTC Val	ACT Thr 2810	8871
GC Al	T GG(a Gly	TG Cy	C CTO	3 TAC u Ty: 281	r Ası	C AGO	C ACC	TG:	r GAT s Asp 2820	Asp	C CG	r GAG	G TTC	C ATO e Met 282	G TGC t Cys 5	8919
CA G1	G AAG n Ası	c cg n Ar	C TT g Le 283	u Cy	T AT	r cco	C AAG b Ly:	G CA' s Hi: 283	s Pho	C GTO	G TG	C GAG	C CA' p Hi: 284	s As	C CGT p Arg	8967

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	TGT Cys 2					Asp					Cys					9015
Cys	GGG Gly 2860				Phe				_	Gly						9063
CGT Arg 2875	CAG Gln	TGG Trp	GAA Glu	Cys	GAT Asp 880	GJ Y GGG	GAG Glu	AAT Asn	Asp	TGT Cys 885	CAC His	GAC Asp	CAC His	Ser	GAT Asp 2890	9111
	GCT Ala		Lys					Thr					Lys			9159
	TCA Ser	Ser					Ser					Val				9207
	CTC Leu					Asp					Gly					9255
Gly	TGC Cys 2940				Glu					Lys						9303
	GAC Asp			Asp					Phe					Arg		9351
	TTC Phe		Leu					Arg					Leu			9399
	AGC Ser	Thr					Ser					Asn		His		9447 .
			Cys			Val					Pro		Gly		GAC Asp	9495
	CAC His 3020	Ser			Ala		Thr			Glu		Phe				9543
	Asn					Arg					Asp				TAC Tyr 3050	9591 -
AC? Thi	A CTG	CTT Leu	AAG Lys	CAG Glr 3055	Gly	CTG	AAC Asr	TAA : Asr	GCG Ala 3060	Val	GCC Ala	teu	GCA Ala	A TTT a Phe 3065	GAC Asp	.96 3 9
TAC Tyl	C CGA	GAG Glu	G CAC Glr 3070	Met	ATC : Ile	TAC Tyr	TGC Trp	ACC Thi 3075	: Gly	C GTO Val	G ACC	C ACC	C CAC Gl: 3080	n Gly	C AGC y Ser	9687

FIG. 12A

(Sheet 34 of 91)

ATG Met	Ile	CGC Arg 085	AGG Arg	ATG Met	CAC His	Leu	AAC Asn 8090	GGC Gly	AGC Ser	AAC Asn	Val	CAG Gln 8095	GTT Val	CTG Leu	CAC His	9735
Arg	ACG Thr 3100	GGC Gly	CTT Leu	AGT Ser	Asn	CCA Pro 105	GAT Asp	GGG Gly	CTC Leu	Ala	GTG Val 3110	GAC Asp	TGG Trp	GTG Val	GGT Gly	9783
GGC Gly 3115	AAC Asn	CTG Leu	TAC Tyr	TGG Trp	TGT Cys 3120	GAC Asp	AAG Lys	GGC Gly	Arg	GAT Asp 125	ACC Thr	ATT Ile	GAG Glu	Val	TCC Ser 3130	9831
AAG Lys	CTT Leu	AAC Asn	Gly	GCC Ala 3135	TAT Tyr	CGG Arg	ACA Thr	Val	CTG Leu 3140	GTC Val	AGC Ser	TCT Ser	Gly	CTC Leu 3145	CGG Arg	9879
GAG Glu	CCC Pro	Arg	GCT Ala 3150	CTG Leu	GTA Val	GTG Val	Asp	GTA Val 3155	CAG Gln	AAT Asn	GGG Gly	Tyr	CTG Leu 3160	TAC Tyr	TGG Trp	9927
ACA Thr	Asp	TGG Trp 3165	GGT Gly	GAC Asp	CAC His	Ser	CTG Leu 3170	ATC Ile	GGC	CGG Arg	Ile	GGC Gly 3175	ATG Met	GAT Asp	GGA Gly	9975
Ser				ATC Ile	Ile					Ile						10023
CTG Leu 3195	ACC Thr	GTG Val	GAC Asp	TAC Tyr	GTC Val 3200	ACG Thr	GAA Glu	CGC Arg	Ile	TAC Tyr 3205	TGG Trp	GCT Ala	GAC Asp	Ala	CGT Arg 3210	10071
			Ile	GAG Glu 3215				Leu					Arg			10119
		Ser		GAC Asp			His					Thr				10167 .
	Tyr		Tyr	TGG Trp		Asp					Ser		Asn		GCC Ala	. 10215
His		Thr		GGT Gly	Ala					Leu					CAC His	10263
	Pro			Leu					Ala		Arg				GTG Val 3290	10311
			Pro		Lys					Gly					TGC Cys	10359
				Gly					Cys					Ası	TTC Phe	10407

FIG. 12A

(Sheet 35 of 91)

TAT Tyr	Leu	GGT Gly 3325	GGC Gly	GAT Asp	GGC Gly	Arg	ACC Thr 330	TGT Cys	GTG Val	TCC Ser	AAC Asn 3	TGC Cys 335	ACA Thr	GCA Ala	AGC Ser	10455
Gln	TTT Phe 3340	GTG Val	TGC Cys	AAA Lys	Asn	GAC Asp 345	AAG Lys	TGC Cys	ATC Ile	Pro	TTC Phe 3350	TGG Trp	TGG Trp	AAG Lys	TGT Cys	10503
GAC Asp 3355	ACG Thr	GAG Glu	GAC Asp	Asp	TGT Cys 360	Gly GGG	GAT Asp	CAC His	Ser	GAC Asp 3365	GAG Glu	CCT Pro	CCA Pro	Asp	TGT Cys 3370	10551
CCC Pro	GAG Glu	TTC Phe	Lys	TGC Cys 3375	CGC Arg	CCA Pro	GGC Gly	Gln	TTC Phe 3380	CAG Gln	TGC Cys	TCC Ser	Thr	GGC Gly 3385	ATC Ile	10599
TGC Cys	ACC Thr	Asn	CCT Pro 3390	GCC Ala	TTC Phe	ATC Ile	Cys	GAT Asp 395	GGG Gly	GAC Asp	AAT Asn	Asp	TGC Cys 8400	CAA Gln	GAC Asp	10647
AAT Asn	Ser	GAC Asp 3405	GAG Glu	GCC Ala	AAT Asn	Cys	GAC Asp 3410	ATT Ile	CAC His	GTC Val	TGC Cys	TTG Leu 8415	CCC Pro	AGC Ser	CAA Gln	10695
Phe	AAG Lys 3420	TGC Cys	ACC Thr	AAC Asn	Thr	AAC Asn 3425	CGC Arg	TGC Cys	ATT Ile	Pro	GGC Gly 3430	ATC Ile	TTC Phe	CGT Arg	TGC Cys	10743
AAT Asn 3435	GGG Gly	CAG Gln	GAC Asp	Asn	TGC Cys 3440	GGG Gly	GAC Asp	GGC Gly	Glu	GAT Asp 3445	GAG Glu	CGG Arg	GAT Asp	Cys	CCT Pro 3450	10791
GAG Glu	GTG Val	ACC Thr	Cys	GCC Ala 3455	CCC Pro	AAC Asn	CAG Gln	Phe	CAG Gln 3460	TGC Cys	TCC Ser	ATC Ile	Thr	AAG Lys 3465	CGC Arg	10839
TGC Cys	ATC Ile	Pro	CGC Arg 3470	GTC Val	TGG Trp	GTC Val	Cys	GAC Asp 3475	AGG Arg	GAT Asp	AAT Asn	His	TGT Cys 3480	GTG Val	GAC Asp	10887
	Ser					Asn									GAT Asp ·	10935
Glu					Asp					Ile	CCC Pro 3510				AAG Lys	10983
				Asp					Gly		GAT Asp					11031
GAG Glu	TGT Cys	GAT Asp	Glu	CGC Arg 3535	ACC Thr	TGT Cys	GAG Glu	Pro	TAC Tyr 3540	Gln	TTC Phe	CGC Arg	Çys	AAA Lys 3545	Asn	11079
		Cys					Trp		Cys			Asp		Asp	TGC Cys	11127

FIG. 12A

(Sheet 36 of 91)

	GAT Asp 3					Glu					Arg					11175
Ser	GAG Glu 3580				Ala					Ile						11223
	GAT Asp			His					Gly					Asp		11271
	CCC Pro		Cys					Phe					Gly			11319
	CCC Pro	Leu					Asp					Cys			_	11367
	GAC Asp					Gly					Thr					11415
Glu	TTT Phe 3660				Asn					Pro						11463
	GGA Gly			Asp					Ser					Glu		11511
	GCC Ala		Phe					Asn					Cys		Asn	11559
	CGA Arg	Val		Leu			Gly					Gly		Asp		11607
TGT Cys	Gly	GAT Asp 3725	Gly	ACT Thr	GAC Asp	Glu	GAG Glu 3730	GAC Asp	TGT Cys	GAG Glu	Pro	CCC Pro 3735	ACG	GCC	CAG Gln	11655
	CCC Pro 3740	His					Lys					Arg				11703
TG1 Cys 3755	Leu	TCA Ser	TCC Ser	TCC Ser	CTG Leu 3760	Arg	TGT Cys	AAC Asn	ATG Met	TTC Phe 3765	a Asp	GAC Asp	TGC Cys	GGG Gly	GAT Asp 3770	11751 · .
GG(C TCC	GAT Asp	GAP Glu	GAA Glu 3775	Asp	TGC Cys	AGC Ser	ATC Ile	GAC Asp 3780	Pro	C AAG D Lys	CTG Leu	Th:	2 AGC 2 Sec 378	C TGT r Cys 5	11799
				s Ser					Glu					l Ar	C ACT g Thr	11847

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04	143	- 13	+										10	116	3L 01	Oi ·	J 1
GAG Glu	Lys	GCT Ala 8805	GCC Ala	TAC Tyr	TGT Cys	Ala	TGC Cys 810	CGC '	TCG Ser	Gly	Phe	CAT His 815	ACT Thr	GTG Val	CCG Pro	11895	
Gly		CCC Pro			Gln					Cys						11943	_
		CAG Gln		Trp					Gly					Ser		11991	
		AAC Asn	Phe					Asn					Glu			12039	
		CAG Gln					Ala					Ile				12087	
	Pro	GGC Gly 3885				Ser					Thr					12135	
Glu		GTC Val			Asp					His						12183	}
	Tyr	TGG Trp		Asn					Thr					Ser		12231	
		GCC Ala	Ala					Ser					Arg			12279	€
		GGT Gly		Thr			Asn					Lys		Pro		1232	7
GGT Gly	ATC	GCT Ala 3965	Ile	GAC Asp	TGG Trp	Val	GCC Ala 3970	Gly GGG	AAT Asn	GTG Val	Tyr	TGG Trp 3975	Thr	GAT Asp	TCC Ser	1237	5
GGC	CGA Arg 3980	Asp	GTG Val	ATT Ile	Glu	GTG Val 3985	GCG Ala	CAA Gln	ATG Met	Lys	GGC Gly 3990	Glu	AAC Asn	CGC Arg	AAG Lys	1242	3
ACC Thi 3995	Lev	ATC	TCG	GGC	ATG Met	Ile	GAT Asp	GAG Glu	CCC	CAT His 4005	Ala	ATC Ile	GTG Val	GTG Val	GAC Asp 4010	1247	1 -
					Met			Şer		Trp					AAG Lys	1251	9
AT:	GAI	A ACA	GCF Ala 4030	Ala	ATC Met	G GAT	Gly	ACC Thr 4035	Let	CGC	G GAG G Glu	AC1	CTC Let 4040	ı Va:	G CAA L Gln	1256	57

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0-1		. •	•										•			
GAC Asp	Asn	ATT Ile 045	CAG Gln	TGG Trp	CCT Pro	Thr	GGG Gly 050	CTG Leu	GCT Ala	GTG Val	Asp	TAT Tyr 1055	CAC His	AAT Asn	GAA Glu	12615
Arg					Asp					Val		GGC Gly				12663
				Asp					Ala			AAA Lys		Gly		12711
			Phe					Phe				ATC Ile	Tyr			12759
		Ile					Phe					TTT Phe				12807
	Leu					Gly					Ala	TCT Ser 4135				12855
Leu					Lys					Thr		CCC Pro				12903
				Trp					Ser			GGG Gly		Val		12951
			Asn					Asp				TGT Cys	Val			12999
		Pro					Asp					GGA Gly				13047
	Gln					Gly		Cys				GCT Ala 4215	Arg		CAG Gln	13095
Pro		Cys					Arg					AAG Lys				13143
	Gln			Glu		Cys					Thi	TGT Cys				13191
					Thr					Thr		C TTC y Phe			, Pro	13,239
				Gln					y Tyi					ı Sei	C ACC	13287

FIG. 12A

(Sheet 39 of 91)

TGC Cys	Thr	GTC Val 285	AAC Asn	CAG Gln	ejà ecc	Asn	CAG Gln 290	CCC Pro	CAG Gln	TGC Cys	Arg	TGT Cys 1295	CTA Leu	CCT Pro	GGC Gly	13335
Phe					Cys					Cys		GGC Gly				13383
				Cys					Asp			CGA Arg		Cys		13431
			Tyr					Arg				AAC Asn	Lys			13479
		Leu					Val					ACC Thr				13527
	Cys					Gly					Ser	TGT Cys 4375				13575
Ile					Asn					Thr		AAC Asn				13623
				Gln					Met			CCC Pro				13671
			Val					Pro				GCC Ala	Ser		Leu	13719
		Leu		Leu			Leu		Leu			GCT Ala		Val		13767 .
TTC Phe	Trp	TAT Tyr 4445	Lys	CGG Arg	CGA Arg	Val	CGA Arg 4450	Gly	GCT Ala	AAG Lys	GGC Gly	TTC Phe 4455	Gln	CAC His	CAG Gln	13815
		Thr			Ala		Asn					Asn	_		TAC Tyr	13863 ∵
	Met					Glu					. Gl				GAT Asp 4490	13911
					Asp					Thi					C CCA n Pro	13959
				Leu					y His					s Se	C CTG r Leu	Ĩ4007

GCC AGC ACG GAC GAG AAG CGA GAA CTG CTG GGC CGG GGA CCT GAA GAC Ala Ser Thr Asp Glu Lys Arg Glu Leu Leu Gly Arg Gly Pro Glu Asp 4535 4525 4530 GAG ATA GGA GAT CCC TTG GCA TAGGGCCCTG CCCCGACGGA TGTCCCCAGA AAGC 14110 CCCCTGCCAC ATGAGTCTTT CAATGAACCC CCTCCCCAGC CGGCCCTTCT CCGGCCCTGC 14170 Glu Ile Gly Asp Pro Leu Ala CGGGTGTACA AATGTAAAAA TGAAGGAATT ACTTTTTATA TGTGAGCGAG CAAGCGAGCA 14230 AGCACAGTAT TATCTCTTTG CATTTCCTTC CTGCCTGCTC CTCAGTATCC CCCCCATGCT GCCTTGAGGG GGCGGGGAGG GCTTTGTGGC TCAAAGGTAT GAAGGAGTCC ACATGTTCCC 14350 TACCGAGCAT ACCCCTGGAA GCCTGGCGGC ACGGCCTCCC CACCACGCCT GTGCAAGACA CTCAACGGG CTCCGTGTCC CAGCTTTCCT TTCCTTGGCT CTCTGGGGTT AGTTCAGGGG 14470 AGGTGGAGTC CTCTGCTGAC CCTGTCTGGA AGATTTGGCT CTAGCTGAGG AAGGAGTCTT TTAGTTGAGG GAAGTCACCC CAAACCCCAG CTCCCACTTT CAGGGGCACC TCTCAGATGG CCATGCTCAG TATCCCTTCC AGACAGGCCC TCCCCTCTCT AGCGCCCCCT CTGTGGCTCC 14650 TAGGGCTGAA CACATTCTTT GGTAACTGTC CCCCAAGCCT CCCATCCCCC TGAGGGCCAG 14710 GAAGAGTCGG GGCACACCAA GGAAGGGCAA GCGGGCAGCC CCATTTTGGG GACGTGAACG 14770
TTTTAATAAT TTTTGCTGAA TTCCTTTACA ACTAAATAAC ACAGATATTG TTATAAATAA 14830 AATTGTAAAA AAAAAAAAA

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1				5	Leu				10					15	
Val	Ser	Gly	Ala 20	Thr	Met	Asp	Ala	Pro 25	Lys	Thr	Cys	Ser	Pro 30	Lys	Gln
Phe	Ala	Cys 35	Arg	Asp	Gln	Ile	Thr 40	Cys	Ile	Ser	Lys	Gly 45	Trp	Arg	Cys
Asp	Gly 50		Arg	Asp	Cys	Pro 55	Asp	Gly	Ser	Asp	Glu 60		Pro	Glu	Ile
Cys 65	Pro	Gln	Ser	Lys	Ala 70	Gln	Arg	Cys	Pro	Pro 75		Glu	His	Ser	Cys 80
Leu	-			85	Cys				90					95	
	_		100		Gly			105					110		
-		115			Arg		120					125			
	130				Cys	135					140				
145	_				Lys 150					155					160
Cys				165	Thr				170					175	
			180		Leu			185					190		
		195			Arg		200					205			
	210				Tyr	215					220				
225	Thr				Gln 230					235					240
Glu				245	Val				250					255	
	-	_	260		Met			265					270		
		275			Leu		280					285			
-	290				Asn	295					300				
305					Arg 310					315					320
Leu	Gļu			325					330					335	
_			340	i	Asp			345					350		
_		355	•		Asn		360					365			
	370)				375	Į.				380				Trp
305					390	1				395					Lys 400
GJ?	Arg			405	•				410)	•			415	
			420)				425	5				430)	Ala
		439	5				44()				445	,		Ser
Thi	Glu 450		c. Glr	ı Val	. Val	Th:	Arq	y Val	l Asp) Lys	Gl ₃	y Gly)	r Ala	a Let	His

FIG. 12B

```
Ile Tyr His Gln Arg Arg Gln Pro Arg Val Arg Ser His Ala Cys Glu
                    470
                                       475
Asn Asp Gln Tyr Gly Lys Pro Gly Gly Cys Ser Asp Ile Cys Leu Leu
                485 .
                                    490
Ala Asn Ser His Lys Ala Arg Thr Cys Arg Cys Arg Ser Gly Phe Ser
                                505
Leu Gly Ser Asp Gly Lys Ser Cys Lys Lys Pro Glu His Glu Leu Phe
                            520
                                              525
Leu Val Tyr Gly Lys Gly Arg Pro Gly Ile Ile Arg Gly Met Asp Met
                        535
                                           540
Gly Ala Lys Val Pro Asp Glu His Met Ile Pro Ile Glu Asn Leu Met
                  550
                                       555
Asn Pro Arg Ala Leu Asp Phe His Ala Glu Thr Gly Phe Ile Tyr Phe
               565
                                    570
                                                       575
Ala Asp Thr Thr Ser Tyr Leu Ile Gly Arg Gln Lys Ile Asp Gly Thr
                                585
            580
Glu Arg Glu Thr Ile Leu Lys Asp Gly Ile His Asn Val Glu Gly Val
                            600
                                                605
Ala Val Asp Trp Met Gly Asp Asn Leu Tyr Trp Thr Asp Asp Gly Pro
                        615
    610
                                            620
Lys Lys Thr Ile Ser Val Ala Arg Leu Glu Lys Ala Ala Gln Thr Arg
                   630
                                        635
Lys Thr Leu Ile Glu Gly Lys Met Thr His Pro Arg Ala Ile Val Val
        . 645
                                    650
Asp Pro Leu Asn Gly Trp Met Tyr Trp Thr Asp Trp Glu Glu Asp Pro
                                665
         660
                                                   670
Lys Asp Ser Arg Arg Gly Arg Leu Glu Arg Ala Trp Met Asp Gly Ser
                             680
                                                685
His Arg Asp Ile Phe Val Thr Ser Lys Thr Val Leu Trp Pro Asn Gly
                        695
    690
Leu Ser Leu Asp Ile Pro Ala Gly Arg Leu Tyr Trp Val Asp Ala Phe
                    710
                                       715
Tyr Asp Arg Ile Glu Thr Ile Leu Leu Asn Gly Thr Asp Arg Lys Ile
                725
                                                        735
                                    730
Val Tyr Glu Gly Pro Glu Leu Asn His Ala Phe Gly Leu Cys His His
            740
                               745
                                                    750
Gly Asn Tyr Leu Phe Trp Thr Glu Tyr Arg Ser Gly Ser Val Tyr Arg
                             760
        755
                                                765
Leu Glu Arg Gly Val Ala Gly Ala Pro Pro Thr Val Thr Leu Leu Arg
                         775
                                            780
Ser Glu Arg Pro Pro Ile Phe Glu Ile Arg Met Tyr Asp Ala His Glu
                    790
                                        795
Gln Gln Val Gly Thr Asn Lys Cys Arg Val Asn Asn Gly Gly Cys Ser
                805
                                    810
Ser Leu Cys Leu Ala Thr Pro Gly Ser Arg Gln Cys Ala Cys Ala Glu
                                                   830
                                825
            820
Asp Gln Val Leu Asp Thr Asp Gly Val Thr Cys Leu Ala Asn Pro Ser
                            840
                                               845
 Tyr Val Pro Pro Pro Gln Cys Gln Pro Gly Gln Phe Ala Cys Ala Asn
                         855
 Asn Arg Cys Ile Gln Glu Arg Trp Lys Cys Asp Gly Asp Asn Asp Cys
 865 . 870
                                        875
 Leu Asp Asn Ser Asp Glu Ala Pro Ala Leu Cys His Gln His Thr Cys
                885
                                     890
                                                        895
 Pro Ser Asp Arg Phe Lys Cys Glu Asn Asn Arg Cys Ile Pro Asn Arg
                                 905
             900
                                                    910
 Trp Leu Cys Asp Gly Asp Asn Asp Cys Gly Asn Ser Glu Asp Glu Ser
                             920
         915
```

FIG. 12B

```
Asn Ala Thr Cys Ser Ala Arg Thr Cys Pro Pro Asn Gln Phe Ser Cys
                 935
                                    940
Ala Ser Gly Arg Cys Ile Pro Ile Ser Trp Thr Cys Asp Leu Asp Asp
                950
                                 955
Asp Cys Gly Asp Arg Ser Asp Glu Ser Ala Ser Cys Ala Tyr Pro Thr
            965
                             970
Cys Phe Pro Leu Thr Gln Phe Thr Cys Asn Asn Gly Arg Cys Ile Asn
                        985
                                          990
Ile Asn Trp Arg Cys Asp Asn Asp Asn Asp Cys Gly Asp Asn Ser Asp
      995 1000
                                       1005
Glu Ala Gly Cys Ser His Ser Cys Ser Ser Thr Gln Phe Lys Cys Asn
                         1020
                  1015
Ser Gly Arg Cys Ile Pro Glu His Trp Thr Cys Asp Gly Asp Asn Asp
               1030
                               1035
Cys Gly Asp Tyr Ser Asp Glu Thr His Ala Asn Cys Thr Asn Gln Ala
                            1050
           1045
                                             1055
Thr Arg Pro Pro Gly Gly Cys His Ser Asp Glu Phe Gln Cys Pro Leu
        1060
                         1065
                                          1070
Asp Gly Leu Cys Ile Pro Leu Arg Trp Arg Cys Asp Gly Asp Thr Asp
  1075 1080
                                     1085
Cys Met Asp Ser Ser Asp Glu Lys Ser Cys Glu Gly Val Thr His Val
          1095
                           1100
Cys Asp Pro Asn Val Lys Phe Gly Cys Lys Asp Ser Ala Arg Cys Ile
     1110 1115
Ser Lys Ala Trp Val Cys Asp Gly Asp Ser Asp Cys Glu Asp Asn Ser
                            1130
            1125
Asp Glu Glu Asn Cys Glu Ala Leu Ala Cys Arg Pro Pro Ser His Pro
                         1145
Cys Ala Asn Asn Thr Ser Val Cys Leu Pro Pro Asp Lys Leu Cys Asp
                      1160
     1155
                                      1165
Gly Lys Asp Asp Cys Gly Asp Gly Ser Asp Glu Gly Glu Leu Cys Asp
                  1175
                                   1180
Gln Cys Ser Leu Asn Asn Gly Gly Cys Ser His Asn Cys Ser Val Ala
       1190 1195
Pro Gly Glu Gly Ile Val Cys Ser Cys Pro Leu Gly Met Glu Leu Gly
            1205
                            1210
Ser Asp Asn His Thr Cys Gln Ile Gln Ser Tyr Cys Ala Lys His Leu
 1220 1225 1230
Lys Cys Ser Gln Lys Cys Asp Gln Asn Lys Phe Ser Val Lys Cys Ser
           1240
                                      1245
Cys Tyr Glu Gly Trp Val Leu Glu Pro Asp Gly Glu Thr Cys Arg Ser
 1250 1255 1260
Leu Asp Pro Phe Lys Leu Phe Ile Ile Phe Ser Asn Arg His Glu Ile
                               1275
               1270
Arg Arg Ile Asp Leu His Lys Gly Asp Tyr Ser Val Leu Val Pro Gly
            1285
                             1290
                                             1295
Leu Arg Asn Thr Ile Ala Leu Asp Phe His Leu Ser Gln Ser Ala Leu
                          1305
        1300
                                          1310
Tyr Trp Thr Asp Ala Val Glu Asp Lys Ile Tyr Arg Gly Lys Leu Leu
     1315
                      1320
                                       1325
Asp Asn Gly Ala Leu Thr Ser Phe Glu Val Val Ile Gln Tyr Gly Leu
                   1335
                                   1340
Ala Thr Pro Glu Gly Leu Ala Val Asp Trp Ile Ala Gly Asn Ile Tyr
               1350
                                1355
                                                 1360
Trp Val Glu Ser Asn Leu Asp Gln Ile Glu Val Ala Lys Leu Asp Gly
            1365
                           1370
                                            1375
Thr Leu Arg Thr Thr Leu Leu Ala Gly Asp Ile Glu His Pro Arg Ala
                          1385
Ile Ala Leu Asp Pro Arg Asp Gly Ile Leu Phe Trp Thr Asp Trp Asp
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1395 1400 1405 Ala Ser Leu Pro Arg Ile Glu Ala Ala Ser Met Ser Gly Ala Gly Arg 1410 1415 1420 Arg Thr Ile His Arg Glu Thr Gly Ser Gly Gly Cys Ala Asn Gly Leu 1430 1435 Thr Val Asp Tyr Leu Glu Lys Arg Ile Leu Trp Ile Asp Ala Arg Ser 1445 1450 1455 Asp Ala Ile Tyr Ser Ala Arg Tyr Asp Gly Ser Gly His Met Glu Val 1460 1465 1470 Leu Arg Gly His Glu Phe Leu Ser His Pro Phe Ala Val Thr Leu Tyr 1475 1480 1485 Gly Gly Glu Val Tyr Trp Thr Asp Trp Arg Thr Asn Thr Leu Ala Lys 1490 1495 1500 Ala Asn Lys Trp Thr Gly His Asn Val Thr Val Val Gln Arg Thr Asn 1510 1515 Thr Gln Pro Phe Asp Leu Gln Val Tyr His Pro Ser Arg Gln Pro Met 1525 1530 Ala Pro Asn Pro Cys Glu Ala Asn Gly Gly Arg Gly Pro Cys Ser His 1540 1545 1550 Leu Cys Leu Ile Asn Tyr Asn Arg Thr Val Ser Trp Ala Cys Pro His 1555 1560 1565 Leu Met Lys Leu His Lys Asp Asn Thr Thr Cys Tyr Glu Phe Lys Lys 1570 1575 1580 Phe Leu Leu Tyr Ala Arg Gln Met Glu Ile Arg Gly Val Asp Leu Asp 1590 1595 Ala Pro Tyr Tyr Asn Tyr Ile Ile Ser Phe Thr Val Pro Asp Ile Asp 1605 1610 1615 Asn Val Thr Val Leu Asp Tyr Asp Ala Arg Glu Gln Arg Val Tyr Trp 1620 1625 1630 Ser Asp Val Arg Thr Gln Ala Ile Lys Arg Ala Phe Ile Asn Gly Thr 1635 1640 1645 Gly Val Glu Thr Val Val Ser Ala Asp Leu Pro Asn Ala His Gly Leu 1655 1660 Ala Val Asp Trp Val Ser Arg Asn Leu Phe Trp Thr Ser Tyr Asp Thr 665 1670 1675 Asn Lys Lys Gln Ile Asn Val Ala Arg Leu Asp Gly Ser Phe Lys Asn 1685 1690 1695 Ala Val Val Gln Gly Leu Glu Gln Pro His Gly Leu Val Val His Pro 1700 1705 1710 Leu Arg Gly Lys Leu Tyr Trp Thr Asp Gly Asp Asn Ile Ser Met Ala 1715 1720 1725 Asn Met Asp Gly Ser Asn His Thr Leu Leu Phe Ser Gly Gln Lys Gly 1735 1740 Pro Val Gly Leu Ala Ile Asp Phe Pro Glu Ser Lys Leu Tyr Trp Ile 1750 1755 Ser Ser Gly Asn His Thr Ile Asn Arg Cys Asn Leu Asp Gly Ser Glu 1765 1770 Leu Glu Val Ile Asp Thr Met Arg Ser Gln Leu Gly Lys Ala Thr Ala 1780 1785 1790 Leu Ala Ile Met Gly Asp Lys Leu Trp Trp Ala Asp Gln Val Ser Glu 1800 1805 Lys Met Gly Thr Cys Asn Lys Ala Asp Gly Ser Gly Ser Val Val Leu 1815 1820 Arg Asn Ser Thr Thr Leu Val Met His Met Lys Val Tyr Asp Glu Ser 1830 1835 Ile Gln Leu Glu His Glu Gly Thr Asn Pro Cys Ser Val Asn Asn Gly 1845 1850 1855 Asp Cys Ser Gln Leu Cys Leu Pro Thr Ser Glu Thr Thr Arg Ser Cys . 1865

Met Cys Thr Ala Gly Tyr Ser Leu Arg Ser Gly Gln Gln Ala Cys Glu 1875 1880 1885 Gly Val Gly Ser Phe Leu Leu Tyr Ser Val His Glu Gly Ile Arg Gly 1895 1900 1890 Ile Pro Leu Asp Pro Asn Asp Lys Ser Asp Ala Leu Val Pro Val Ser 1920 1910 1915 Gly Thr Ser Leu Ala Val Gly Ile Asp Phe His Ala Glu Asn Asp Thr 1925 1930 1935 Ile Tyr Trp Val Asp Met Gly Leu Ser Thr Ile Ser Arg Ala Lys Arg 1950 1940 1945 Asp Gln Thr Trp Arg Glu Asp Val Val Thr Asn Gly Ile Gly Arg Val 1955 1960 1965 Glu Gly Ile Ala Val Asp Trp Ile Ala Gly Asn Ile Tyr Trp Thr Asp 1970 1975 1980 Gln Gly Phe Asp Val Ile Glu Val Ala Arg Leu Asn Gly Ser Phe Arg 1990 1995 985 Tyr Val Val Ile Ser Gln Gly Leu Asp Lys Pro Arg Ala Ile Thr Val 2005 2010 His Pro Glu Lys Gly Tyr Leu Phe Trp Thr Glu Trp Gly His Tyr Pro 2025 2020 Arg Ile Glu Arg Ser Arg Leu Asp Gly Thr Glu Arg Val Val Leu Val 2045 2040 Asn Val Ser Ile Ser Trp Pro Asn Gly Ile Ser Val Asp Tyr Gln Gly 2060 2055 Gly Lys Leu Tyr Trp Cys Asp Ala Arg Met Asp Lys Ile Glu Arg Ile 2070 2075 Asp Leu Glu Thr Gly Glu Asn Arg Glu Val Val Leu Ser Ser Asn Asn 2095 2090 2085 Met Asp Met Phe Ser Val Ser Val Phe Glu Asp Phe Ile Tyr Trp Ser 2100 2105 2110 Asp Arg Thr His Ala Asn Gly Ser Ile Lys Arg Gly Cys Lys Asp Asn 2120 2125 2115 Ala Thr Asp Ser Val Pro Leu Arg Thr Gly Ile Gly Val Gln Leu Lys 2140 2130 2135 Asp Ile Lys Val Phe Asn Arg Asp Arg Gln Lys Gly Thr Asn Val Cys 2160 2150 2155 Ala Val Ala Asn Gly Gly Cys Gln Gln Leu Cys Leu Tyr Arg Gly Gly 2175 2165 2170 Gly Gln Arg Ala Cys Ala Cys Ala His Gly Met Leu Ala Glu Asp Gly 2185 2190 2180 Ala Ser Cys Arg Glu Tyr Ala Gly Tyr Leu Leu Tyr Ser Glu Arg Thr 2205 2200 Ile Leu Lys Ser Ile His Leu Ser Asp Glu Arg Asn Leu Asn Ala Pro 2215 2220 Val Gln Pro Phe Glu Asp Pro Glu His Met Lys Asn Val Ile Ala Leu 2230 2235 Ala Phe Asp Tyr Arg Ala Gly Thr Ser Pro Gly Thr Pro Asn Arg Ile 2250 2245 Phe Phe Ser Asp Ile His Phe Gly Asn Ile Gln Gln Ile Asn Asp Asp 2270 2260 2265 Gly Ser Gly Arg Thr Thr Ile Val Glu Asn Val Gly Ser Val Glu Gly -2280 2275 Leu Ala Tyr His Arg Gly Trp Asp Thr Leu Tyr Trp Thr Ser Tyr Thr 2295 2300 Thr Ser Thr Ile Thr Arg His Thr Val Asp Gln Thr Arg Pro Gly Ala 2310 2315 2320 Phe Glu Arg Glu Thr Val Ile Thr Met Ser Gly Asp Asp His Pro Arg 2330 2325 Ala Phe Val Leu Asp Glu Cys Gln Asn Leu Met Phe Trp Thr Asn Trp

2340 2345 Asn Glu Leu His Pro Ser Ile Met Arg Ala Ala Leu Ser Gly Ala Asn 2360 2365 Val Leu Thr Leu Ile Glu Lys Asp Ile Arg Thr Pro Asn Gly Leu Ala 2370 2375 2380 Ile Asp His Arg Ala Glu Lys Leu Tyr Phe Ser Asp Ala Thr Leu Asp 385 2390 2395 2400 Lys Ile Glu Arg Cys Glu Tyr Asp Gly Ser His Arg Tyr Val Ile Leu 2405 2410 Lys Ser Glu Pro Val His Pro Phe Gly Leu Ala Val Tyr Gly Glu His 2420 2425 2430 lle Phe Trp Thr Asp Trp Val Arg Arg Ala Val Gln Arg Ala Asn Lys 2435 2440 2445Tyr Val Gly Ser Asp Met Lys Leu Leu Arg Val Asp Ile Pro Gln Gln 2455 2460 Pro Met Gly Ile Ile Ala Val Ala Asn Asp Thr Asn Ser Cys Glu Leu 465 2470 2475 Ser Pro Cys Arg Ile Asn Asn Gly Gly Cys Gln Asp Leu Cys Leu Leu 2485 2490 2495 Thr His Gln Gly His Val Asn Cys Ser Cys Arg Gly Gly Arg Ile Leu 2500 2505 Gln Glu Asp Phe Thr Cys Arg Ala Val Asn Ser Ser Cys Arg Ala Gln 2520 2525 Asp Glu Phe Glu Cys Ala Asn Gly Glu Cys Ile Ser Phe Ser Leu Thr 2530 2535 2540 Cys Asp Gly Val Ser His Cys Lys Asp Lys Ser Asp Glu Lys Pro Ser 545 2550 2555 2560 Tyr Cys Asn Ser Arg Arg Cys Lys Lys Thr Phe Arg Gln Cys Asn Asn 2565 2570 Gly Arg Cys Val Ser Asn Met Leu Trp Cys Asn Gly Val Asp Tyr Cys 2585 2590 Gly Asp Gly Ser Asp Glu Ile Pro Cys Asn Lys Thr Ala Cys Gly Val 2595 2600 2605 Gly Glu Phe Arg Cys Arg Asp Gly Ser Cys Ile Gly Asn Ser Ser Arg 2615 2620 Cys Asn Gln Phe Val Asp Cys Glu Asp Ala Ser Asp Glu Met Asn Cys 625 2630 2635 Ser Ala Thr Asp Cys Ser Ser Tyr Phe Arg Leu Gly Val Lys Gly Val 2645 2650 Leu Phe Gln Pro Cys Glu Arg Thr Ser Leu Cys Tyr Ala Pro Ser Trp 2660 2665 Val Cys Asp Gly Ala Asn Asp Cys Gly Asp Tyr Ser Asp Glu Arg Asp 2675 2680 2685 Cys Pro Gly Val Lys Arg Pro Arg Cys Pro Leu Asn Tyr Phe Ala Cys 2695 2700 Pro Ser Gly Arg Cys Ile Pro Met Ser Trp Thr Cys Asp Lys Glu Asp 2710 2715 Asp Cys Glu Asn Gly Glu Asp Glu Thr His Cys Asn Lys Phe Cys Ser 2725 2730 2735 Glu Ala Gln Phe Glu Cys Gln Asn His Arg Cys Ile Ser Lys Gln Trp 2740 2745 2750 Leu Cys Asp Gly Ser Asp Asp Cys Gly Asp Gly Ser Asp Glu Ala Ala 2755 2760 2765 His Cys Glu Gly Lys Thr Cys Gly Pro Ser Ser Phe Ser Cys Pro Gly . 2775 2780 Thr His Val Cys Val Pro Glu Arg Trp Leu Cys Asp Gly Asp Lys Asp 2790 2795 Cys Thr Asp Gly Ala Asp Glu Ser Val Thr Ala Gly Cys Leu Tyr Asn 2810

Ser Thr Cys Asp Asp Arg Glu Phe Met Cys Gln Asn Arg Leu Cys Ile 2820 2825 Pro Lys His Phe Val Cys Asp His Asp Arg Asp Cys Ala Asp Gly Ser 2840 2845 2835 Asp Glu Ser Pro Glu Cys Glu Tyr Pro Thr Cys Gly Pro Asn Glu Phe 2850 2855 2860 Arg Cys Ala Asn Gly Arg Cys Leu Ser Ser Arg Gln Trp Glu Cys Asp 865 2870 2875 2880 Gly Glu Asn Asp Cys His Asp His Ser Asp Glu Ala Pro Lys Asn Pro 2885 2890 2895 His Cys Thr Ser Pro Glu His Lys Cys Asn Ala Ser Ser Gln Phe Leu 2905 2900 2910 Cys Ser Ser Gly Arg Cys Val Ala Glu Ala Leu Leu Cys Asn Gly Gln 2915 2920 2925 Asp Asp Cys Gly Asp Gly Ser Asp Glu Arg Gly Cys His Val Asn Glu 2930 2935 2940 Cys Leu Ser Arg Lys Leu Ser Gly Cys Ser Gln Asp Cys Glu Asp Leu 2950 2955 Lys Ile Gly Phe Lys Cys Arg Cys Arg Pro Gly Phe Arg Leu Lys Asp 2965 2970 2975 Asp Gly Arg Thr Cys Ala Asp Leu Asp Glu Cys Ser Thr Thr Phe Pro 2980 2985 2990 Cys Ser Gln Leu Cys Ile Asn Thr His Gly Ser Tyr Lys Cys Leu Cys 2995 3000 3005 Val Glu Gly Tyr Ala Pro Arg Gly Gly Asp Pro His Ser Cys Lys Ala 3015 3020 Val Thr Asp Glu Glu Pro Phe Leu Ile Phe Ala Asn Arg Tyr Tyr Leu 3030 3035 Arg Lys Leu Asn Leu Asp Gly Ser Asn Tyr Thr Leu Leu Lys Gln Gly 3050 3055 3045 Leu Asn Asn Ala Val Ala Leu Ala Phe Asp Tyr Arg Glu Gln Met Ile 3060 3065 3070 Tyr Trp Thr Gly Val Thr Thr Gln Gly Ser Met Ile Arg Arg Met His 3085 3075 3080 Leu Asn Gly Ser Asn Val Gln Val Leu His Arg Thr Gly Leu Ser Asn 3090 3095 3100 Pro Asp Gly Leu Ala Val Asp Trp Val Gly Gly Asn Leu Tyr Trp Cys 3110 3115 3120 Asp Lys Gly Arg Asp Thr Ile Glu Val Ser Lys Leu Asn Gly Ala Tyr 3130 3125 Arg Thr Val Leu Val Ser Ser Gly Leu Arg Glu Pro Arg Ala Leu Val 3140 3145 3150 Val Asp Val Gln Asn Gly Tyr Leu Tyr Trp Thr Asp Trp Gly Asp His 3155 3160 3165 Ser Leu Ile Gly Arg Ile Gly Met Asp Gly Ser Gly Arg Ser Ile Ile 3170 3180 Val Asp Thr Lys Ile Thr Trp Pro Asn Gly Leu Thr Val Asp Tyr Val 3190 3195 Thr Glu Arg Ile Tyr Trp Ala Asp Ala Arg Glu Asp Tyr Ile Glu Phe 3205 3210 3215 Ala Ser Leu Asp Gly Ser Asn Arg His Val Val Leu Ser Gln Asp Ile 3220 3225 3230 Pro His Ile Phe Ala Leu Thr Leu Phe Glu Asp Tyr Val Tyr Trp Thr 3240 3245 . 3235 Asp Trp Glu Thr Lys Ser Ile Asn Arg Ala His Lys Thr Thr Gly Ala 3260 3255 Asn Lys Thr Leu Leu Ile Ser Thr Leu His Arg Pro Met Asp Leu His 3275 3280 3270 Val Phe His Ala Leu Arg Gln Pro Asp Val Pro Asn His Pro Cys Lys

3285 3290 Val Asn Asn Gly Gly Cys Ser Asn Leu Cys Leu Leu Ser Pro Gly Gly 3300 3305 3310 Gly His Lys Cys Ala Cys Pro Thr Asn Phe Tyr Leu Gly Gly Asp Gly 3315 3320 3325 Arg Thr Cys Val Ser Asn Cys Thr Ala Ser Gln Phe Val Cys Lys Asn 3335 3340 Asp Lys Cys Ile Pro Phe Trp Trp Lys Cys Asp Thr Glu Asp Asp Cys 3350 3355 3360 Gly Asp His Ser Asp Glu Pro Pro Asp Cys Pro Glu Phe Lys Cys Arg 3365 3370 3375 Pro Gly Gln Phe Gln Cys Ser Thr Gly Ile Cys Thr Asn Pro Ala Phe 3380 3385 3390 Ile Cys Asp Gly Asp Asn Asp Cys Gln Asp Asn Ser Asp Glu Ala Asn 3395 3400 3405Cys Asp Ile His Val Cys Leu Pro Ser Gln Phe Lys Cys Thr Asn Thr 3410 3415 3420 Asn Arg Cys Ile Pro Gly Ile Phe Arg Cys Asn Gly Gln Asp Asn Cys 3430 3435 Gly Asp Gly Glu Asp Glu Arg Asp Cys Pro Glu Val Thr Cys Ala Pro 3445 3450 3455 Asn Gln Phe Gln Cys Ser Ile Thr Lys Arg Cys Ile Pro Arg Val Trp 3465 3470 3460 Val Cys Asp Arg Asp Asn His Cys Val Asp Gly Ser Asp Glu Pro Ala 3475 3480 3485 3475 3480 3485 Asn Cys Thr Gln Met Thr Cys Gly Val Asp Glu Phe Arg Cys Lys Asp 3495 3500 3490 Ser Gly Arg Cys Ile Pro Ala Arg Trp Lys Cys Asp Gly Glu Asp Asp 505 3510 3515 3520 Cys Gly Asp Gly Ser Asp Glu Pro Lys Glu Glu Cys Asp Glu Arg Thr 3525 3530 3535 Cys Glu Pro Tyr Gln Phe Arg Cys Lys Asn Asn Arg Cys Val Pro Gly 3540 3545 Arg Trp Gln Cys Asp Tyr Asp Asn Asp Cys Gly Asp Asn Ser Asp Glu 3555 3560 3565 Glu Ser Cys Thr Pro Arg Pro Cys Ser Glu Ser Glu Phe Phe Cys Ala . 3580 3575 Asn Gly Arg Cys Ile Ala Gly Arg Trp Lys Cys Asp Gly Asp His Asp 585 3590 3595 3600 Cys Ala Asp Gly Ser Asp Glu Lys Asp Cys Thr Pro Arg Cys Asp Met 3605 3610 3615 Asp Gln Phe Gln Cys Lys Ser Gly His Cys Ile Pro Leu Arg Trp Pro 3625 3630 Cys Asp Ala Asp Ala Asp Cys Met Asp Gly Ser Asp Glu Glu Ala Cys 3640 Gly Thr Gly Val Arg Thr Cys Pro Leu Asp Glu Phe Gln Cys Asn Asn 3650 3655 3660 Thr Leu Cys Lys Pro Leu Ala Trp Lys Cys Asp Gly Glu Asp Asp Cys 3675 3670 Gly Asp Asn Ser Asp Glu Asn Pro Glu Glu Cys Ala Arg Phe Ile Cys 3685 3690 Pro Pro Asn Arg Pro Phe Arg Cys Lys Asn Asp Arg Val Cys Leu Trp 3700 3705 3710 Ile Gly Arg Gln Cys Asp Gly Val Asp Asn Cys Gly Asp Gly Thr Asp 3715 3720 3725 3720 3725 Glu Glu Asp Cys Glu Pro Pro Thr Ala Gln Asn Pro His Cys Lys Asp 3735 3740 Lys Lys Glu Phe Leu Cys Arg Asn Gln Arg Cys Leu Ser Ser Ser Leu 3750 3755

Arg Cys Asn Met Phe Asp Asp Cys Gly Asp Gly Ser Asp Glu Glu Asp 3765 3770 Cys Ser Ile Asp Pro Lys Leu Thr Ser Cys Ala Thr Asn Ala Ser Met 3780 3785 3790 Cys Gly Asp Glu Ala Arg Cys Val Arg Thr Glu Lys Ala Ala Tyr Cys 3795 3800 3805 Ala Cys Arg Ser Gly Phe His Thr Val Pro Gly Gln Pro Gly Cys Gln 3815 3820 Asp Ile Asn Glu Cys Leu Arg Phe Gly Thr Cys Ser Gln Leu Trp Asn 3830 3835 Lys Pro Lys Gly Gly His Leu Cys Ser Cys Ala Arg Asn Phe Met Lys 3845 3850 Thr His Asn Thr Cys Lys Ala Glu Gly Ser Glu Tyr Gln Val Leu Tyr 3860 3865 3870 Ile Ala Asp Asp Asn Glu Ile Arg Ser Leu Phe Pro Gly His Pro His 3875 3880 3885 Ser Ala Tyr Glu Gln Thr Phe Gln Gly Asp Glu Ser Val Arg Ile Asp 3895 3900 Ala Met Asp Val His Val Lys Ala Gly Arg Val Tyr Trp Thr Asn Trp 3910 3915 3920 His Thr Gly Thr Ile Ser Tyr Arg Ser Leu Pro Pro Ala Ala Pro Pro 3925 3930 3935 Thr Thr Ser Asn Arg His Arg Arg Gln Ile Asp Arg Gly Val Thr His 3940 3945 3950Leu Asn Ile Ser Gly Leu Lys Met Pro Arg Gly Ile Ala Ile Asp Trp 3955 3960 3965 Val Ala Gly Asn Val Tyr Trp Thr Asp Ser Gly Arg Asp Val Ile Glu 3975 **Val Ala Gln Met Lys Gly Glu Asn Arg Lys Thr Leu Ile Ser Gly Met** 3990 3995 Ile Asp Glu Pro His Ala Ile Val Val Asp Pro Leu Arg Gly Thr Met 4010 4015 4005 Tyr Trp Ser Asp Trp Gly Asn His Pro Lys Ile Glu Thr Ala Ala Met 4020 4025 4030 Asp Gly Thr Leu Arg Glu Thr Leu Val Gln Asp Asn Ile Gln Trp Pro 4040 4045 Thr Gly Leu Ala Val Asp Tyr His Asn Glu Arg Leu Tyr Trp Ala Asp 4050 4055 4060 Ala Lys Leu Ser Val Ile Gly Ser Ile Arg Leu Asn Gly Thr Asp Pro 4070 4075 Ile Val Ala Ala Asp Ser Lys Arg Gly Leu Ser His Pro Phe Ser Ile 4085 4090 Asp Val Phe Glu Asp Tyr Ile Tyr Gly Val Thr Tyr Ile Asn Asn Arg 4100 4105 4110 Val Phe Lys Ile His Lys Phe Gly His Ser Pro Leu Tyr Asn Leu Thr 4115 4120 4125 Gly Gly Leu Ser His Ala Ser Asp Val Val Leu Tyr His Gln His Lys 4135 4140 4130 Gln Pro Glu Val Thr Asn Pro Cys Asp Arg Lys Lys Cys Glu Trp Leu 4150 4155 Cys Leu Leu Ser Pro Ser Gly Pro Val Cys Thr Cys Pro Asn Gly Lys . 4165 4170 4175 Arg Leu Asp Asn Gly Thr Cys Val Pro Val Pro Ser Pro Thr Pro Pro 4185 4190 4180 Pro Asp Ala Pro Arg Pro Gly Thr Cys Thr Leu Gln Cys Phe Asn Gly 4200 4205 4195 Gly Ser Cys Phe Leu Asn Ala Arg Arg Gln Pro Lys Cys Arg Cys Gln 4215 4220 Pro Arg Tyr Thr Gly Asp Lys Cys Glu Leu Asp Gln Cys Trp Glu Tyr

4230 4235 Cys His Asn Gly Gly Thr Cys Ala Ala Ser Pro Ser Gly Met Pro Thr 4245 4250 4255 Cys Arg Cys Pro Thr Gly Phe Thr Gly Pro Lys Cys Thr Ala Gln Val 4260 4265 4270 4265 4270 Cys Ala Gly Tyr Cys Ser Asn Asn Ser Thr Cys Thr Val Asn Gln Gly 4285 4275 4280 Asn Gln Pro Gln Cys Arg Cys Leu Pro Gly Phe Leu Gly Asp Arg Cys 4295 4300 Gln Tyr Arg Gln Cys Ser Gly Phe Cys Glu Asn Phe Gly Thr Cys Gln 305 4310 4315 4320 Met Ala Ala Asp Gly Ser Arg Gln Cys Arg Cys Thr Val Tyr Phe Glu 4325 4330 4335 Gly Pro Arg Cys Glu Val Asn Lys Cys Ser Arg Cys Leu Gln Gly Ala 4340 4345 4350 Cys Val Val Asn Lys Gln Thr Gly Asp Val Thr Cys Asn Cys Thr Asp 4360 4365 4355 Gly Arg Val Ala Pro Ser Cys Leu Thr Cys Ile Asp His Cys Ser Asn 4370 4380 Gly Gly Ser Cys Thr Met Asn Ser Lys Met Met Pro Glu Cys Gln Cys 4390 4395 Pro Pro His Met Thr Gly Pro Arg Cys Gln Glu Gln Val Val Ser Gln 4405 4410 4415 Gln Gln Pro Gly His Met Ala Ser Ile Leu Ile Pro Leu Leu Leu 4420 4425 4430 Leu Leu Leu Leu Val Ala Gly Val Val Phe Trp Tyr Lys Arg Arg 4435 4440 4445 Val Arg Gly Ala Lys Gly Phe Gln His Gln Arg Met Thr Asn Gly Ala 4455 4460 Met Asn Val Glu Ile Gly Asn Pro Thr Tyr Lys Met Tyr Glu Gly Gly 465 4470 4475 4480 Glu Pro Asp Asp Val Gly Gly Leu Leu Asp Ala Asp Phe Ala Leu Asp 4485 4490 Pro Asp Lys Pro Thr Asn Phe Thr Asn Pro Val Tyr Ala Thr Leu Tyr 4500 4505 4510 Met Gly Gly His Gly Ser Arg His Ser Leu Ala Ser Thr Asp Glu Lys 4515 4520 4525 Arg Glu Leu Leu Gly Arg Gly Pro Glu Asp Glu Ile Gly Asp Pro Leu 4535 4530 Ala 545

GCTA	CAATO	CC A	rctgo	GTCT	СТС	CCAGO	CTCC	TTC	TTTC	rgc <i>i</i>	AAC 1		GG A				55
AAA Lys 5	CTC Leu	CTT Leu	CAT His	CCA Pro	AGT Ser 10	CTG Leu	GTT Val	CTT Leu	CTC Leu	CTC Leu 15	TTG Leu	GTC Val	CTC Leu	CTG Leu	CCC Pro 20		103
ACA Thr	GAC Asp	GCC Ala	TCA Ser	GTC Val 25	TCT Ser	GGA Gly	AAA Lys	CCG Pro	CAG Gln 30	TAT Tyr	ATG Met	GTT Val	CTG Leu	GTC Val 35	CCC Pro	:	151
TCC Ser	CTG Leu	CTC Leu	CAC His 40	ACT Thr	GAG Glu	ACC Thr	ACT Thr	GAG Glu 45	AAG Lys	GGC Gly	TGT Cys	GTC Val	CTT Leu 50	CTG Leu	AGC Ser	:	199
TAC Tyr	CTG Leu	AAT Asn 55	GAG Glu	ACA Thr	GTG Val	ACT Thr	GTA Val 60	AGT Ser	GCT Ala	TCC Ser	TTG Leu	GAG Glu 65	TCT Ser	GTC Val	AGG Arg	:	247
GGA Gly	AAC Asn 70	AGG Arg	AGC Ser	CTC Leu	TTC Phe	ACT Thr 75	GAC Asp	CTG Leu	GAG Glu	GCG Ala	GAG Glu 80	AAT Asn	GAC Asp	GTA Val	CTC Leu	;	295
CAC His 85	TGT Cys	GTC Val	GCC Ala	TTC Phe	GCT Ala 90	GTC Val	CCA Pro	AAG Lys	TCT Ser	TCA Ser 95	TCC Ser	AAT Asn	GAG Glu	GAG Glu	GTA Val 100	:	343
ATG Met	TTC Phe	CTC Leu	ACT Thr	GTC Val 105	CAA Gln	GTG Val	AAA Lys	GGA Gly	CCA Pro 110	ACC Thr	CAA Gln	GAA Glu	TTT Phe	AAG Lys 115	AAG Lys	:	391
CGG Arg	ACC Thr	ACA Thr	GTG Val 120	ATG Met	GTT Val	AAG Lys	AAC Asn	GAG Glu 125	GAC Asp	AGT Ser	CTG Leu	GTC Val	TTT Phe 130	GTC Val	CAG Gln	•	439
ACA Thr	GAC Asp	AAA Lys 135	TCA Ser	ATC Ile	TAC Tyr	AAA Lys	CCA Pro 140	GGG Gly	CAG Gln	ACA Thr	GTG Val	AAA Lys 145	TTT Phe	CGT Arg	GTT Val	•	487
GTC Val	TCC Ser 150	ATG Met	GAT Asp	GAA Glu	AAC Asn	TTT Phe 155	CAC His	CCC Pro	CTG Leu	AAT Asn	GAG Glu 160	TTG Leu	ATT Ile	CCA Pro	CTA Leu	!	535
GTA Val 165	TAC Tyr	ATT Ile	CAG Gln	GAT Asp	CCC Pro 170	AĄA Lys	GGA Gly	AAT Asn	CGC Arg	ATC Ile 175	GCA Ala	CAA Gln	TGG Trp	CAG Gln	AGT Ser 180	!	583
TTC Phe	CAG Gln	TTA Leu	GAG Glu	GGT Gly 185	GGC Gly	CTC Leu	AAG Lys	CAA Gln	TTT Phe 190	TCT Ser	TTT Phe	Pro	CTC Leu	TCA Ser 195	TCA Ser	-	631
GAG Glu	CCC Pro	TTC Phe	CAG Gln 200	GGC Gly	TCC Ser	TAC Tyr	AAG Lys	GTG Val 205	GTG Val	GTA Val	CAG Gln	AAG Lys	ÁAA Lys 210	TCA Ser	GGT Gly		679
GGA	AGG	ACA	GAG	CAC	ССТ	TTC	ACC	GTG	GAG	GAA	TTT	GTT	СТТ	ССС	AAG		727

FIG. 13A

(Sheet 52 of 91)

Gly	Arg	Thr 215	Glu	His	Pro	Phe	Thr 220	Val	Glu	Glu	Phe	Val 225	Leu	Pro	Lys	
TTT Phe	GAA Glu 230	GTA Val	CAA Gln	GTA Val	ACA Thr	GTG Val 235	CCA Pro	AAG Lys	ATA Ile	ATC Ile	ACC Thr 240	ATC Ile	TTG Leu	GAA Glu	GAA Glu	775
GAG Glu 245	ATG Met	AAT Asn	GTA Val	TCA Ser	GTG Val 250	TGT Cys	GGC Gly	CTA Leu	TAC Tyr	ACA Thr 255	TAT Tyr	GGG Gly	AAG Lys	CCT Pro	GTC Val 260	823
												AGT Ser				871
GAC Asp	TGC Cys	CAC His	GGT Gly 280	GAA Glu	GAT Asp	TCA Ser	CAG Gl n	GCT Ala 285	TTC Phe	TGT Cys	GAG Glu	AAA Lys	TTC Phe 290	AGT Ser	GGA Gly	919
												AAA Lys 305				967
												CAC His				1015
												GGA Gly				1063
												GTG Val				1111
									Phe			GTG Val				1159
								Asn				TTC Phe 385				1207
												GAG Glu				1255
	Gln										Gly	ACC Thr				1303
					Lys					Cys		GGC Gly			Trp	1351 ·
				His					His					. Val	TTC Phe	1399

FIG. 13A

(Sheet 53 of 91)

						CTT Leu			Met				1447
						CAG Gln							1495
						CTC Leu							1543
 						GJA GGG							1591
						TCC Ser 525							1639
						CTC Leu							1687
						AAA Lys							1735
						AGC Ser							1783
						GCG Ala							1831
						CTG Leu 605							1879
		Ser				Leu						ACT . Thr	1927
	Pro				Asp	CAG Gln				Asp			1975
His						ATC Ile			Thr				2023
				Met		TTC Phe		Glu				Lys	2071
			Ser				Pro				Glr	CTT Leu	2119

FIG. 13A

(Sheet 54 of 91)

CAA Gln	CAG Gln	TAT Tyr 695	GAA Glu	ATG Met	CAT His	GGA Gly	CCT Pro 700	GAA Glu	GGT Gly	CTA Leu	CGT Arg	GTA Val 705	GGT Gly	TTT Phe	TAT Tyr	2167
GAG Glu	TCA Ser 710	GAT Asp	GTA Val	ATG Met	GGA Gly	AGA Arg 715	GGC GGC	CAT His	GCA Ala	CGC Arg	CTG Leu 720	GTG Val	CAT His	GTT Val	GAA Glu	2215
				GAG Glu												2263
				GTG Val 745												2311
ACA Thr	GTC Val	CCT Pro	GAC Asp 760	ACC Thr	ATC Ile	ACC Thr	GAG Glu	TGG Trp 765	AAG Lys	GCA Ala	GGG Gly	GCC Ala	TTC Phe 770	TGC Cys	CTG Leu	2359
				GGA Gly												2407
				TTT Phe												2455
				ACA Thr												2503
				AGT Ser 825												2551
				AAG Lys												2599
				TCC Ser				Thr								2647
		Thr		AGC Ser			Ala					Glu				2695
ACT Thr 885	Glu	GTG Val	CCT Pro	TCA Ser	GTT Val 890	Pro	GAA Glu	. CAC . His	GGA Gly	AGG Arg 895	Lys	GAC Asp	ACA Thr	GTC Val	ATC Ile 900	2743
				GTT Val 905	Glu					Glu					Phe	2791
				Cys					/ Glu					Let	TCC Ser	2839

FIG. 13A

8449-134 (Sheet 55 of 91)

U	4-4-4) - I 4	J-T										, -			
CTG Leu	AAA Lys	CTG Leu 935	CCA Pro	CCA Pro	AAT Asn	GTG Val	GTA Val 940	GAA Glu	GAA Glu	TCT Ser	GCC Ala	CGA Arg 945	GCT Ala	TCT Ser	GTC Val	2887
					ATA Ile											2935
					TAT Tyr 970											2983
					GTA Val											3031
		Glu			TCC Ser		Ala					Asn				3079
	Arg				TAC Tyr	Lys					Ser					3127
Gly					AGG Arg					Thr						3175
				Phe	GCC Ala 1050				Ala					Asp		3223
			Thr		GCC Ala			Trp		Ser					Asp	3271
		Cys			AGC Ser		Gly		Leu			Asn		Ile		3319
	Gly				GAA Glu	Val		Leu					Thr			3367
Leu		Glu			Leu		Val					Val			GCC Ala	3415
	Phe					Ala					Glr				CAT His 1140	3463
					Thr					ı Ala					C CTG a Leu 5	3511
				Asp					ı Val					u Ası	r GAG n Glu	3559

FIG. 13A

(Sheet 56 of 91)

GAA Glu	GCT Ala 1	GTG Val 175	AAG Lys	AAA Lys	GAC Asp	Asn	TCT Ser 180	GTC Val	CAT His	TGG Trp	Glu	CGC Arg 185	CCT Pro	CAG Gln	AAA Lys	3607
Pro	AAG Lys 190	GCA Ala	CCA Pro	GTG Val	Gly	CAT His 195	TTT Phe	TAC Tyr	GAA Glu	Pro	CAG Gln 200	GCT Ala	CCC Pro	TCT Ser	GCT Ala	3655
	GTG Val			Thr					Leu					Ala		3703
	GCC Ala		Thr					Thr					Ile			3751
	ATC Ile	Thr					Ala					Ser				3799
	ACA Thr					His					Tyr					3847
Phe	ACC Thr 1270				Lys					Thr						3895
	TTT Phe			Lys					Asn					Leu		3943
	CAG Gln		Ser					Pro					Met			3991
	GGA Gly	Glu					Leu					Lys		Asn		4039
	CCA Pro		Lys			Phe		Phe			Gly		Gln			4087
	CAA Gln 1350				Glu		Lys					Phe				4135
	AGT Ser			Tyr		Gly			Ser		Ser					4183
	GAT Asp				Val					Pro					Val	4231
				Arg					. Ser					L Se	C AGC c Ser	4279

FIG. 13A

(Sheet 57 of 91)

AAC	CAT	GTC	TTG	ATT	TAC	CTT	GAT	AAG	GTG	TCA	AAT	CAG	ACA	CTG	AGC	4327	
Asn			Leu	Ile	Tyr			Lys	Val	Ser	Asn	Gln	Thr	Leu	Ser		
	1	L415					1420				:	1425					
			ACG													4375	
Leu	Phe	Phe	Thr	Val	Leu	Gln	Asp	Val	Pro	Val	Arg	Asp	Leu	Lys	Pro		
3	430				1	1435					144Ó			_			
GCC	ATA	GTG	AAA	GTC	TAT	GAT	TAC	TAC	GAG	ACG	GAT	GAG	TTT	GCA	ATC	4423	
Ala	Ile	Val	Lvs	Val	Tyr	Asp	Tyr	Tvr	Glu	Thr	Asp	Glu	Phe	Ala	Ile		
1445					1450	-	-	_		1455	-				1460		
									-					•			
GCT	GAG	TAC	AAT	GCT	CCT	TGC	AGC	AAA	GAT	CTT	GGA	AAT	GCT	TGA	AGACCA	4474	
			Asn														
		-1-		1465		-1-		_	1470	200	-			7			
			•					•						_			
CAAC	SCCTO	LAAF	AAGTO	CTT	rg C	rgga	GTCC	r GT	rere	CAC	CTC	CACA	CAD (GACA	CGTGTT	4534	
			TAAA									O. LOFE	·		001011	4577	
***	3 7 2 7 7 6		* * T-71.7/	J J		11	******	J 240.		1010	GIC					2311	

```
Ser Val Ser Gly Lys Pro Gln Tyr Met Val Leu Val Pro Ser Leu Leu
His Thr Glu Thr Thr Glu Lys Gly Cys Val Leu Leu Ser Tyr Leu Asn
           20
                                25
Glu Thr Val Thr Val Ser Ala Ser Leu Glu Ser Val Arg Gly Asn Arg
                           40
Ser Leu Phe Thr Asp Leu Glu Ala Glu Asn Asp Val Leu His Cys Val
Ala Phe Ala Val Pro Lys Ser Ser Ser Asn Glu Glu Val Met Phe Leu
                    70
                                       75
Thr Val Gln Val Lys Gly Pro Thr Gln Glu Phe Lys Lys Arg Thr Thr
               85
                                    90
Val Met Val Lys Asn Glu Asp Ser Leu Val Phe Val Gln Thr Asp Lys
            100
                                105
                                                    110
Ser Ile Tyr Lys Pro Gly Gln Thr Val Lys Phe Arg Val Val Ser Met
                           120
        115
                                               125
Asp Glu Asn Phe His Pro Leu Asn Glu Leu Ile Pro Leu Val Tyr Ile
                       135
                                          140
Gln Asp Pro Lys Gly Asn Arg Ile Ala Gln Trp Gln Ser Phe Gln Leu
                                       155
                    150
Glu Gly Gly Leu Lys Gln Phe Ser Phe Pro Leu Ser Ser Glu Pro Phe
                                    170
Gln Gly Ser Tyr Lys Val Val Gln Lys Lys Ser Gly Gly Arg Thr
            180
                                185
Glu His Pro Phe Thr Val Glu Glu Phe Val Leu Pro Lys Phe Glu Val
                            200
                                               205
Gln Val Thr Val Pro Lys Ile Ile Thr Ile Leu Glu Glu Met Asn
                         215
                                            220
Val Ser Val Cys Gly Leu Tyr Thr Tyr Gly Lys Pro Val Pro Gly His
                    230
                                        235
Val Thr Val Ser Ile Cys Arg Lys Tyr Ser Asp Ala Ser Asp Cys His
                245
                                    250
Gly Glu Asp Ser Gln Ala Phe Cys Glu Lys Phe Ser Gly Gln Leu Asn
                                265
                                                    270
Ser His Gly Cys Phe Tyr Gln Gln Val Lys Thr Lys Val Phe Gln Leu
                                                285
                            280
        275
Lys Arg Lys Glu Tyr Glu Met Lys Leu His Thr Glu Ala Gln Ile Gln
                         295
                                            300
Glu Glu Gly Thr Val Val Glu Leu Thr Gly Arg Gln Ser Ser Glu Ile
                                        315
                     310
Thr Arg Thr Ile Thr Lys Leu Ser Phe Val Lys Val Asp Ser His Phe
                                    330
                 325
 Arg Gln Gly Ile Pro Phe Phe Gly Gln Val Arg Leu Val Asp Gly Lys
                                345
 Gly Val Pro Ile Pro Asn Lys Val Ile Phe Ile Arg Gly Asn Glu Ala
                             360
         355
 Asn Tyr Tyr Ser Asn Ala Thr Thr Asp Glu His Gly Leu Val Gln Phe
                         375
                                            380
 Ser Ile Asn Thr Thr Asn Val Met Gly Thr Ser Leu Thr Val Arg Val
                     390
                                         395
 Asn Tyr Lys Asp Arg Ser Pro Cys Tyr Gly Tyr Gln Trp Val Ser Glu
                 405
                                    .410
 Glu His Glu Glu Ala His His Thr Ala Tyr Leu Val Phe Ser Pro Ser
                                                     430
             420
                                 425
 Lys Ser Phe Val His Leu Glu Pro Met Ser His Glu Leu Pro Cys Gly
                            440
                                                 445
         435
 His Thr Gln Thr Val Gln Ala His Tyr Ile Leu Asn Gly Gly Thr Leu
                                          460
                         455
 Leu Gly Leu Lys Lys Leu Ser Phe Tyr Tyr Leu Ile Met Ala Lys Gly
```

FIG. 13B

(Sheet 59 of 91)

													•		
465					470					475					480
				400	Gly				490					105	
Met	Lys	Gly	His 500	Phe	Ser	Ile	Ser	Ile 505	Pro	Val	Lys	Ser	Asp 510	Ile	Ala
Pro	Val	Ala 515	Arg	Leu	Leu	Ile	Tyr 520	Ala	Val	Leu	Pro		Gly	Asp	Val
Ile	Gly 530		Ser	Ala	Lys	Tyr 535	Asp	Val	Glu	Asn		525 Leu	Ala	Asn	Lys
Val		Leu	Ser	Phe	Ser		Ser	Gln	Ser		540 Pro	Ala	Ser	His	Ala
545 His	Leu	Arg	Val	Thr 565	550 Ala	Ala	Pro	Gln	Ser	555 Val	Cys	Ala	Leu		560 Ala
Val	Asp	Gln	Ser 580		Leu	Leu	Met	Lys 585	570 Pro	Asp	Ala	Glu		575 Ser	Ala
Ser	Ser	Val 595		Asn	Leu	Leu	Pro 600		Lys	Asp	Leu		590 Gly	Phe	Pro
Gly	Pro 610		Asn	Asp	Gln	Asp 615	Asp	Glu	Asp	Cys	Ile 620	605 Asn	Arg	His	Asn
Val 625	Tyr	Ile	Asn	Gly	Ile 630		Tyr	Thr	Pro	Val 635	Ser	Ser	Thr	Asn	Glu 640
				645	Phe				650	Gly				655	Thr
			660		Lys			665					670		
		6/5			Glu		680					685			
	690				His	695					700				
705					Lys 710					715					720
				725	Ala				730					735	
			740		Trp			745					750		
		755			Ser		760					765			
	770				Thr	775					780	-	_		
785					Thr 790					795					800
				805	Glu -				810					815	
			820		Pro			825					830		
		835			Thr		840					845			
	850				Leu	855					860				
862					His 870					875					880
				885	Gly				890					895	
			900		Gly			905					910		
		915	_		Glu		920					925			
erA	930	тте	Leu	GTÀ	Ser	935	met	GIn	Asn	Thr	Gln 940	Asn	Leu	Leu	Gln

FIG. 13B

```
Met Pro Tyr Gly Cys Gly Glu Gln Asn Met Val Leu Phe Ala Pro Asn
                                        955
                    950
Ile Tyr Val Leu Asp Tyr Leu Asn Glu Thr Gln Gln Leu Thr Pro Glu
               965
                                  970
Val Lys Ser Lys Ala Ile Gly Tyr Leu Asn Thr Gly Tyr Gln Arg Gln
                              985
           980
Leu Asn Tyr Lys His Tyr Asp Gly Ser Tyr Ser Thr Phe Gly Glu Arg
995 1000 1005
Tyr Gly Arg Asn Gln Gly Asn Thr Trp Leu Thr Ala Phe Val Leu Lys
                       1015
                                          1020
Thr Phe Ala Gln Ala Arg Ala Tyr Ile Phe Ile Asp Glu Ala His Ile
                                                          1040
                                      1035
                  1030
Thr Gln Ala Leu Ile Trp Leu Ser Gln Arg Gln Lys Asp Asn Gly Cys
               1045 1050 1055
Phe Arg Ser Ser Gly Ser Leu Leu Asn Asn Ala Ile Lys Gly Gly Val
                 1065 1070
          1060
Glu Asp Glu Val Thr Leu Ser Ala Tyr Ile Thr Ile Ala Leu Leu Glu
                 1080 1085
     1075
Ile Pro Leu Thr Val Thr His Pro Val Val Arg Asn Ala Leu Phe Cys
  1090 1095
                              1100
Leu Glu Ser Ala Trp Lys Thr Ala Gln Glu Gly Asp His Gly Ser His
                   1110 1115
Val Tyr Thr Lys Ala Leu Leu Ala Tyr Ala Phe Ala Leu Ala Gly Asn
                                  1130
         1125
                                                      1135
Gln Asp Lys Arg Lys Glu Val Leu Lys Ser Leu Asn Glu Glu Ala Val
           1140 1145
Lys Lys Asp Asn Ser Val His Trp Glu Arg Pro Gln Lys Pro Lys Ala
                                   1165
                           1160
  1155
Pro Val Gly His Phe Tyr Glu Pro Gln Ala Pro Ser Ala Glu Val Glu
                       1175 1180
Met Thr Ser Tyr Val Leu Leu Ala Tyr Leu Thr Ala Gln Pro Ala Pro
                                      1195
                 1190
Thr Ser Glu Asp Leu Thr Ser Ala Thr Asn Ile Val Lys Trp Ile Thr
               1205
                                   1210 1215
 Lys Gln Gln Asn Ala Gln Gly Gly Phe Ser Ser Thr Gln Asp Thr Val
                   1225 1230
           1220
 Val Ala Leu His Ala Leu Ser Lys Tyr Gly Ala Ala Thr Phe Thr Arg
1235 1240 1245
 Thr Gly Lys Ala Ala Gln Val Thr Ile Gln Ser Ser Gly Thr Phe Ser
                                           1260
   1250
                        1255
 Ser Lys Phe Gln Val Asp Asn Asn Asn Arg Leu Leu Gln Gln Val
                                                           1280
                    1270
                                        1275
 Ser Leu Pro Glu Leu Pro Gly Glu Tyr Ser Met Lys Val Thr Gly Glu
                                    1290
                                                       1295
                1285
 Gly Cys Val Tyr Leu Gln Thr Ser Leu Lys Tyr Asn Ile Leu Pro Glu
 1300 1305 1310

Lys Glu Glu Phe Pro Phe Ala Leu Gly Wals Gln Thr Leu Pro Gln Thr
1315 1320 1325
                                                    1310
 Lys Glu Giu Pne Florin 1320 1325
1315 1320 1325
Cys Asp Glu Pro Lys Ala His Thrisers Phe Gin Tie Ser Leu Ser Wal
1335 1340
 1330
Ser Tyr Thr Gly Ser Arg Ser Ala Ser Ash Met Ala Tie Val Asp Val
 Ser Arg Ser Arg Ser Arg Ser Ara Ser Asn Met Alaille Val Asp val 1355 1360

Lys Met Val Ser Gly Phe Ile Pro Leu Lys Pro Thr Val Lys Met Leu 1365 1370 1375

Glu Arg Ser Asn His Val Ser Arg Thr Glu Val Ser Ser Asn His Val 1380 1385 1390

Leu Ile Tyr Leu Asp Lys Val Ser Asn Gln Thr Leu Ser Leu Phe Phe 1395 1400 1405
```

FIG. 13B

Thr -Val Leu Gln Asp Val Pro Val Arg Asp Leu Lys Pro Ala le Val 1410 1415 1420
Lys Val Tyr Asp Tyr Tyr Glu Thr Asp Glu Phe Ala le Ala Glu Tyr 425 1430 1435 1440
Asn Ala Pro Cys Ser Lys Asp Leu Gly Asn Ala 1445 1445

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CAGCGGTGCG AGCTCCAGGC CCATGCACTG AGGAGGCGGA AACAAGGGGA GCCCCCAGAG CTCCATCAAG CCCCCTCCAA AGGCTCCCCT ACCCGGTCCA CGCCCCCAC CCCCCTCCC CGCCTCCTCC CAATTGTGCA TTTTTGCAGC CGGAGGCGGC TCCGAGATGG GGCTGTGAGC TTCGCCCGGG GAGGGGGAAA GAGCAGCGAG GAGTGAAGCG GGGGGTGGG GTGAAGGGTT TGGATTTCGG GGCAGGGGC GCACCCCCGT CAGCAGGCCC TCCCCAAGGG GCTCGGAACT CTACCTCTTC ACCCACGCCC CTGGTGCGCT TTGCCGAAGG AAAGAATAAG AACAGAGAAG GAGGAGGGGG AAAGGAAGAA AAGGGGGACC CCCCAACTGG GGGGGTGAA GGAGAGAAGT AGCAGGACCA GAGGGGAAGG GGCTGCTGCT TGCATCAGCC CACACC ATG CTG ACC Met Leu Thr 1 CCG CCG TTG CTC CTG CTG CCC CTG CTC TCA GCT CTG GTC GCG GCG Pro Pro Leu Leu Leu Leu Pro Leu Leu Ser Ala Leu Val Ala Ala														
GCT ATC GA Ala Ile As 20						Ala Cys .								
GAT CAA A' Asp Gln I														
GAC TGC CO Asp Cys P					Ile Cys									
AAG GCC CA Lys Ala G	ln Arg Cys													
CTG TGT G Leu Cys V 85														
GAC GGC TO Asp Gly S 100														
TCT CGC C Ser Arg L		Gln His												
ACC TGC T Thr Cys T	AC TGC AAG yr Cys Asi 135			Leu Gln	Ala Asp									
Cys Lys A	AT TTT GAT sp Phe Asp 50													
	AC ACA GAG		Phe Ile											
	CAG CCG GA				Lys Asn									

FIG. 14A

(Sheet 63 of 91)

GAC Asp	CGG Arg	CCC Pro	CCT Pro	GTG Val 200	CTG Leu	TTG Leu	ATA Ile	GCC Ala	AAC Asn 205	TCC Ser	CAG Gln	AAC Asn	ATC Ile	TTG Leu 210	GCC Ala	1099
ACG Thr	TAC Tyr	CTG Leu	AGT Ser 215	GGG Gly	GCC Ala	CAG Gln	GTG Val	TCT Ser 220	ACC Thr	ATC Ile	ACA Thr	CCT Pro	ACG Thr 225	AGC Ser	ACG Thr	1147
CGG Arg	CAG Gln	ACC Thr 230	ACA Thr	GCC Ala	ATG Met	GAC Asp	TTC Phe 235	AGC Ser	TAT Tyr	GCC Ala	AAC Asn	GAG Glu 240	ACC Thr	GTA Val	TGC Cys	1195
TGG Trp	GTG Val 245	CAT His	GTT Val	GGG Gly	GAC Asp	AGT Ser 250	GCT Ala	GCT Ala	CAG Gln	ACG Thr	CAG Gln 255	CTC Leu	AAG Lys	TGT Cys	GCC Ala	1243
CGC Arg 260	ATG Met	CCT Pro	Gly GGC	CTA Leu	AAG Lys 265	GGC Gly	TTC Phe	GTG Val	GAT Asp	GAG Glu 270	CAC His	ACC Thr	ATC Ile	AAC Asn	ATC Ile 275	1291
TCC Ser	CTC Leu	AGT Ser	CTG Leu	CAC His 280	CAC His	GTG Val	GAA Glu	CAG Gln	ATG Met 285	GCC Ala	ATC Ile	GAC Asp	TGG Trp	CTG Leu 290	ACA Thr	1339
GGC Gly	AAC Asn	TTC Phe	TAC Tyr 295	TTT Phe	GTG Val	GAT Asp	GAC Asp	ATC Ile 300	GAT Asp	GAT Asp	AGG Arg	ATC Ile	TTT Phe 305	GTC Val	TGC Cys	1387
AAC Asn	AGA Arg	AAT Asn 310	GGG Gly	GAC Asp	ACA Thr	TGT Cys	GTC Val 315	ACA Thr	TTG Leu	CTA Leu	GAC Asp	CTG Leu 320	GAA Glu	CTC Leu	TAC Tyr	1435
AAC Asn	CCC Pro 325	AAG Lys	GGC Gly	ATT Ile	GCC Ala	CTG Leu 330	GAC Asp	CCT Pro	GCC Ala	ATG Met	GGG Gly 335	AAG Lys	GTG Val	TTT Phe	TTC Phe	1483
ACT Thr 340	GAC Asp	TAT Tyr	GGG Gly	CAG Gln	ATC Ile 345	CCA Pro	AAG Lys	GTG Val	Glu	CGC Arg 350	TGT Cys	GAC Asp	ATG Met	GAT Asp	GGG Gly 355	1531 .
CAG Gln	AAC Asn	CGC Arg	ACC Thr	AAG Lys 360	CTC Leu	GTC Val	GAC Asp	AGC Ser	AAG Lys 365	ATT Ile	GTG Val	TTT Phe	CCT Pro	CAT His 370	GGC Gly ·	1579
ATC Ile	ACG Thr	CTG Leu	GAC Asp 375	CTG Leu	GTC Val	AGC Ser	CGC Arg	CTT Leu 380	GTC Val	TAC Tyr	TGG Trp	GCA Ala	GAT Asp 385	GCC Ala	TAT Tyr	1627
CTG Leu	GAC Asp	TAT Tyr 390	ATT Ile	GAA Glu	GTG Val	GTG Val	GAC Asp 395	TAT Tyr	GAG Glu	GGC Gly	AAG Lys	GGC Gly 400	CGC Arg	CAG Gln	ACC Thr	1675
ATC Ile	ATC Ile 405	CAG Gln	GGC Gly	ATC Ile	CTG Leu	ATT Ile 410	GAG Glu	CAC His	CTG Leu	TAC Tyr	GGC Gly 415	CTG Leu	ACT Thr	GTG Val	TTT Phe	1723
GAG Glu 420	AAT Asn	TAT Tyr	CTC Leu	TAT Tyr	GCC Ala 425	ACC Thr	AAC Asn	TCG Ser	GAC Asp	AAT Asn 430	GCC Ala	AAT Asn	GCC Ala	CAG Gln	CAG Gln 435	1771

FIG. 14A

(Sheet 64 of 91)

AAG ACG AGT (Lys Thr Ser \	GTG ATC CGT Val Ile Arg 440	GTG AAC CGC Val Asn Arg	TTT AAC AGC Phe Asn Ser 445	ACC GAG TAC CAG Thr Glu Tyr Gln 450	1819
Val Val Thr	CGG GTG GAC Arg Val Asp 455	AAG GGT GGT Lys Gly Gly 460	Ala Leu His	ATC TAC CAC CAG Ile Tyr His Gln 465	1867
AGG CGT CAG Arg Arg Gln 470	CCC CGA GTG Pro Arg Val	AGG AGC CAT Arg Ser His 475	GCC TGT GAA Ala Cys Glu	AAC GAC CAG TAT Asn Asp Gln Tyr 480	1915
GGG AAG CCG Gly Lys Pro 485	GGT GGC TGC Gly Gly Cys	TCT GAC ATC Ser Asp Ile 490	TGC CTG CTG Cys Leu Leu 495	GCC AAC AGC CAC Ala Asn Ser His	1963 s
AAG GCG CGG Lys Ala Arg 500	ACC TGC CGC Thr Cys Arg 505	Cys Arg Ser	GGC TTC AGC Gly Phe Ser 510	CTG GGC AGT GAG Leu Gly Ser Asj 51	
GGG AAG TCA Gly Lys Ser	TGC AAG AAC Cys Lys Lys 520	CCG GAG CAT Pro Glu His	GAG CTG TTC Glu Leu Phe 525	C CTC GTG TAT GG e Leu Val Tyr Gl 530	C 20 59 y
AAG GGC CGG Lys Gly Arg	CCA GGC ATO Pro Gly Ilo 535	C ATC CGG GGG E Ile Arg Gly 540	y Met Asp Met	G GGG GCC AAG GT t Gly Ala Lys Va 545	C 2107 1
CCG GAT GAG Pro Asp Glu 550	His Met Il	C CCC ATT GA e Pro Ile Gl 555	A AAC CTC ATo u Asn Leu Me	G AAC CCC CGA GC t Asn Pro Arg Al 560	C 2155 a
CTG GAC TTC Leu Asp Phe 565	CAC GCT GA His Ala Gl	G ACC GGC TT u Thr Gly Ph 570	C ATC TAC TT e Ile Tyr Ph 57	T GCC GAC ACC AC e Ala Asp Thr Th 5	cc 2203 ir
AGC TAC CTC Ser Tyr Leu 580	: ATT GGC CG Ile Gly Ar 58	g Gln Lys Il	T GAT GGC AC e Asp Gly Th 590	T GAG CGG GAG AC Tr Glu Arg Glu Th 59	cc 2251 . nr 95
Ile Leu Lys	Asp Gly II 600	e His Asn Va.	al Glu Gly Va 605	G GCC GTG GAC TO al Ala Val Asp T: 610	rp .
ATG GGA GAC	AAT CTG TA Asn Leu Ty 615	r Trp Thr As	AC GAT GGG CO sp Asp Gly Pi 20	CC AAA AAG ACA A ro Lys Lys Thr I 625	
AGC GTG GCC Ser Val Ala 63	a Arg Leu G	AG AAA GCT G lu Lys Ala A 635	CT CAG ACC CO la Gln Thr A	GC AAG ACT TTA A rg Lys Thr Leu I 640	
GAG GGC AA Glu Gly Ly 645	A ATG ACA C s Met Thr H	AC CCC AGG G is Pro Arg A 650	Ta lle Agt A	TG GAT CCA CTC A al Asp Pro Leu A 55	1311
GGG TGG AT Gly Trp Me 660	t Tyr Trp T	CA GAC TGG G hr Asp Trp G 65	GAG GAG GAC C Glu Glu Asp P 670	CC AAG GAC AGT (Pro Lys Asp Ser A	CGG 2491 Arg 675

FIG. 14A

(Sheet 65 of 91)

CGT Arg	GGG Gly	CGG Arg	CTG Leu	GAG Glu 680	AGG Arg	GCG Ala	TGG Trp	ATG Met	GAT Asp 685	GGC Gly	TCA Ser	CAC His	CGA Arg	GAC Asp 690	ATC Ile	2539
TTT Phe	GTC Val	ACC Thr	TCC Ser 695	AAG Lys	ACA Thr	GTG Val	CTT Leu	TGG Trp 700	CCC Pro	AAT Asn	GGG Gly	CTA Leu	AGC Ser 705	CTG Leu	GAC Asp	2587
ATC Ile	CCG Pro	GCT Ala 710	GGG Gly	CGC Arg	CTC Leu	TAC Tyr	TGG Trp 715	GTG Val	GAT Asp	GCC Ala	TTC Phe	TAC Tyr 720	GAC Asp	CGC Arg	ATC Ile	2635
GAG Glu	ACG Thr 725	ATA Ile	CTG Leu	CTC Leu	AAT Asn	GGC Gly 730	ACA Thr	GAC Asp	CGG Arg	AAG Lys	ATT Ile 735	GTG Val	TAT Tyr	GAA Glu	GGT Gly	2683
CCT Pro 740	GAG Glu	CTG Leu	AAC Asn	CAC His	GCC Ala 745	TTT Phe	GGC	CTG Leu	TGT Cys	CAC His 750	CAT His	GGC Gly	AAC Asn	TAC Tyr	CTC Leu 755	2731
TTC Phe	TGG Trp	ACT Thr	GAG Glu	TAT Tyr 760	CGG Arg	AGT Ser	GGC Gly	AGT Ser	GTC Val 765	TAC Tyr	CGC Arg	TTG Leu	GAA Glu	CGG Arg 770	GGT Gly	2779
GTA Val	GGA Gly	GGC Gly	GCA Ala 775	CCC Pro	CCC Pro	ACT Thr	GTG Val	ACC Thr 780	CTT Leu	CTG Leu	CGC Arg	AGT Ser	GAG Glu 785	CGG Arg	CCC Pro	2827
CCC Pro	ATC Ile	TTT Phe 790	GAG Glu	ATC Ile	CGA Arg	ATG Met	TAT Tyr 795	GAT Asp	GCC Ala	CAG Gln	Gln	CAG Gln 800	CAA Gln	GTT Val	GGC	2875
ACC Thr	AAC Asn 805	AAA Lys	TGC Cys	CGG Arg	GTG Val	AAC Asn 810	AAT Asn	GGC Gly	GGC Gly	TGC Cys	AGC Ser 815	AGC Ser	CTG Leu	TGC Cys	TTG Leu	2923
GCC Ala 820	ACC Thr	CCT Pro	GGG Gly	AGC Ser	CGC Arg 825	CAG Gln	TGC Cys	GCC Ala	TGT Cys	GCT Ala 830	GAG Glu	GAC Asp	CAG Gln	GTG Val	TTG Leu 835	2971 .
GAC Asp	GCA Ala	GAC Asp	GGC Gly	GTC Val 840	ACT Thr	TGC Cys	TTG Leu	GCG Ala	AAC Asn 845	CCA Pro	TCC Ser	TAC Tyr	GTG Val	CCT Pro 850	CCA Pro .	3019
							TTT Phe									3067
CAG Gln	GAG Glu	CGC Arg 870	TGG Trp	AAG Lys	TGT Cys	GAC Asp	GGA Gly 875	GAC Asp	AAC Asn	GAT Asp	TGC Cys	CTG Leu 880	GAC Asp	AAC Asn	AGT Ser	3115
GAT Asp	GAG Glu 885	GCC Ala	CCA Pro	GCC Ala	CTC Leu	TGC Cys 890	CAT	CAG Gln	CAC His	ACC Thr	TGC Cys 895	CCC Pro	TCG Ser	GAC Asp	CGA Arg	3163
TTC Phe 900	Lys	TGC Cys	GAG Glu	AAC Asn	AAC Asn 905	CGG Arg	TGC Cys	ATC Ile	CCC Pro	AAC Asn 910	CGC Arg	TGG Trp	CTC Leu	TGC Cys	GAC Asp 915	3211

FIG. 14A

8449-134 (Sheet 66 of 91) GGG GAC AAT GAC TGT GGG AAC AGT GAA GAT GAG TCC AAT GCC ACT TGT 3259 Gly Asp Asn Asp Cys Gly Asn Ser Glu Asp Glu Ser Asn Ala Thr Cys 920 TCA GCC CGC ACC TGC CCC CCC AAC CAG TTC TCC TGT GCC AGT GGC CGC 3307 Ser Ala Arg Thr Cys Pro Pro Asn Gln Phe Ser Cys Ala Ser Gly Arg 940 945 TGC ATC CCC ATC TCC TGG ACG TGT GAT CTG GAT GAC GAC TGT GGG GAC 3355 Cys Ile Pro Ile Ser Trp Thr Cys Asp Leu Asp Asp Cys Gly Asp 955 CGC TCT GAT GAG TCT GCT TCG TGT GCC TAT CCC ACC TGC TTC CCC CTG 3403 Arg Ser Asp Glu Ser Ala Ser Cys Ala Tyr Pro Thr Cys Phe Pro Leu ACT CAG TIT ACC TGC AAC AAT GGC AGA TGT ATC AAC ATC AAC TGG AGA 3451 Thr Gln Phe Thr Cys Asn Asn Gly Arg Cys Ile Asn Ile Asn Trp Arg 990 TGC GAC AAT GAC AAT GAC TGT GGG GAC AAC AGT GAC GAA GCC GGC TGC 3499 Cys Asp Asn Asp Asn Asp Cys Gly Asp Asn Ser Asp Glu Ala Gly Cys 1005 AGC CAC TCC TGT TCT AGC ACC CAG TTC AAG TGC AAC AGC GGG CGT TGC 3547 Ser His Ser Cys Ser Ser Thr Gln Phe Lys Cys Asn Ser Gly Arg Cys 1015 1020 1025 ATC CCC GAG CAC TGG ACC TGC GAT GGG GAC AAT GAC TGC GGA GAC TAC 3595 Ile Pro Glu His Trp Thr Cys Asp Gly Asp Asn Asp Cys Gly Asp Tyr 1030 1035 AGT GAT GAG ACA CAC GCC AAC TGC ACC AAC CAG GCC ACG AGG CCC CCT 3643 Ser Asp Glu Thr His Ala Asn Cys Thr Asn Gln Ala Thr Arg Pro Pro 1045 1050 GGT GGC TGC CAC ACT GAT GAG TTC CAG TGC CGG CTG GAT GGA CTA TGC 3691 . Gly Gly Cys His Thr Asp Glu Phe Gln Cys Arg Leu Asp Gly Leu Cys 1065 1070 1075 ATC CCC CTG CGG TGG CGC TGC GAT GGG GAC ACT GAC TGC ATG GAC TCC 3739 Ile Pro Leu Arg Trp Arg Cys Asp Gly Asp Thr Asp Cys Met Asp Ser 1085 AGC GAT GAG AAG AGC TGT GAG GGA GTG ACC CAC GTC TGC GAT CCC AGT 3787 Ser Asp Glu Lys Ser Cys Glu Gly Val Thr His Val Cys Asp Pro Ser 1095 1100 GTC AAG TTT GGC TGC AAG GAC TCA GCT CGG TGC ATC AGC AAA GCG TGG 3835 Val Lys Phe Gly Cys Lys Asp Ser Ala Arg Cys Ile Ser Lys Ala Trp 1115 1120 GTG TGT GAT GGC GAC AAT GAC TGT GAG GAT AAC TCG GAC GAG GAG AAC 3883 Val Cys Asp Gly Asp Asn Asp Cys Glu Asp Asn Ser Asp Glu Glu Asn 1125 1130 1135 TGC GAG TCC CTG GCC TGC AGG CCA CCC TCG CAC CCT TGT GCC AAC AAC 3931 Cys Glu Ser Leu Ala Cys Arg Pro Pro Ser His Pro Cys Ala Asn Asn

FIG. 14A

1150

1145

(Sheet 67 of 91)

ACC Thr	TCA Ser	GTC Val	Cys	CTG Leu 160	CCC Pro	CCT Pro	GAC Asp	Lys	CTG Leu 165	TGT Cys	GAT Asp	GGC Gly	Asn	GAC Asp 170	GAC Asp	3979
TGT Cys	GGC	Asp	GGC Gly 175	TCA Ser	GAT Asp	GAG Glu	Gly	GAG Glu 180	CTC Leu	TGC Cys	GAC Asp	Gln	TGC Cys 185	TCT Ser	CTG Leu	4027
AAT Asn	Asn	GGT Gly 190	GGC Gly	TGC Cys	AGC Ser	His	AAC Asn 195	TGC Cys	TCA Ser	GTG Val	GCA Ala 1	CCT Pro 200	GGC Gly	GAA Glu	GGC Gly	4075
Ile					Pro					Leu	GGG Gly .215					4123
				Gln					Lys		CTC Leu			Ser		4171
			Gln					Val			TCC Ser		Tyr			4219
		Leu					Glu				AGC Ser	Leu				4267
	Pro					Ser					ATC Ile					4315
Leu					Tyr					Pro	GGC Gly 1295					4363
				Phe					Ser		CTC Leu			Thr		4411 ·
		-	Asp					Gly		Leu	CTG Leu		Asn		Ala	4459
		Ser					Ile		Tyr		CTG Leu	Ala				4507
	Leu					Ile		Gly			TAC Tyr		Val			4555 -
Asn		Asp			Glu		Ala			Asp	GGG Gly 1375				ACC Thr	4603
	Leu					Ile					Ala				GAT Asp 1395	4651

FIG. 14A

8449-134 (Sheet 68 of 91)

													-			
CCC Pro	CGG Arg	GAT Asp	Gly	ATC Ile 400	CTG Leu	TTT Phe	TGG Trp	Thr	GAC Asp 405	TGG Trp	GAT Asp	GCC Ala	Ser	CTG Leu 410	CCC Pro	4699
		Glu				ATG Met	Ser					Arg				4747
	Glu					GGC Gly 1					Leu					4795
Leu					Leu	TGG Trp 450				Arg						4843
				Asp		TCT Ser			Met					Gly		4891
			Ser			TTT Phe		Val					Gly			4939
		Thr				ACA Thr	Asn					Ala				4987
	Gly					GTG Val					Asn					5035
Asp					His	CCC Pro 1530				Pro						5083
	Glu			Gly		CAG Gln			Cys					Leu		5131
AAC Asn	TAC Tyr	AAC Asn	Arg	ACC Thr 1560	Val	TCC Ser	TGC Cys	Ala	TGC Cys 1565	CCC Pro	CAC His	CTC Leu	Met	AAG Lys 1570	Leu	5179
CAC His	AAG Lys	Asp	AAC Asn 1575	Thr	ACC Thr	TGC Cys	Tyr	GAG Glu 1580	Phe	AAG Lys	AAG Lys	Phe	CTG Leu 1585	Leu	TAC	5227
	Arg		Met			CGA Arg		Val			Asp		Pro			5275 -
		Ile					Val					Asn			GTG Val	5323
	Asp					Glu					Trp				G CGG L Arg 1635	5371

FIG. 14A

(Sheet 69 of 91)

ACA Thr	CAG Gln	GCC Ala	rre	AAG Lys 1640	CGG Arg	GCC Ala	TTC Phe	Ile	AAC Asn 1645	GGC Gly	ACA Thr	GGC Gly	Val	GAG Glu 1650	ACA Thr	5419
GTC Val	GTC Val	Ser	GCA Ala 1655	GAC Asp	TTG Leu	CCA Pro	Asn	GCC Ala 1660	CAC His	GGG Gly	CTG Leu	Ala	GTG Val 665	GAC Asp	TGG Trp	5467
GTC Val	TCC Ser	CGA Arg 1670	AAC Asn	CTG Leu	TTC Phe	Trp	ACA Thr 1675	AGC Ser	TAT Tyr	GAC Asp	Thr	AAT Asn 680	AAG Lys	AAG Lys	CAG Gln	5515
ile	AAT Asn 1685	GTG Val	GCC Ala	CGG Arg	Leu	GAT Asp 1690	GGC Gly	TCC Ser	TTC Phe	Lys	AAC Asn 695	GCA Ala	GTG V al	GTG Val	CAG Gln	5563
GGC Gly 1700	CTG Leu	GAG Glu	CAG Gln	Pro	CAT His 1705	GGC Gly	CTT Leu	GTC Val	Val	CAC His 1710	CCT Pro	CTG Leu	CGT Arg	Gly	AAG Lys 1715	5611
CTC Leu	TAC Tyr	TGG Trp	Thr	GAT Asp .720	GGT Gly	GAC Asp	AAC Asn	Ile	AGC Ser 1725	ATG Met	GCC Ala	AAC Asn	Met	GAT Asp 1730	GGC Gly	5659
AGC Ser	AAT Asn	Arg	ACC Thr 1735	CTG Leu	CTC Leu	TTC Phe	Ser	GGC Gly 1740	CAG Gln	AAG Lys	Gly	Pro	GTG Val 745	GGC Gly	CTG Leu	5707
GCT Ala	ATT Ile	GAC Asp 1750	TTC Phe	CCT Pro	GAA Glu	Ser	AAA Lys .755	CTC Leu	TAC Tyr	TGG Trp	Ile	AGC Ser 760	TCC Ser	GG Gly	AAC Asn	5755
His	ACC Thr 1765	ATC Ile	AAC Asn	CGC Arg	Cys	AAC Asn .770	CTG Leu	GAT Asp	GGG Gly	Ser	GGG Gly 775	CTG Leu	GAG Glu	GTC Val	ATC Ile	5803
GAT Asp 1780	GCC Ala	ATG Met	CGG Arg	Ser	CAG Gln L785	CTG Leu	GGC Gly	AAG Lys	Ala	ACC Thr 790	GCC Ala	CTG Leu	GCC Ala	Ile	ATG Met L795	5851
GGG Gly	GAC Asp	AAG Lys	Leu	TGG Trp .800	TGG Trp	GCT Ala	GAT Asp	Gln	GTG Val 1805	TCG Ser	GAA Glu	AAG Lys	Met	GGC Gly 1810	ACA Thr	5899
TGC Cys	AGC Ser	Lys	GCT Ala 1815	GAC Asp	GGC Gly	TCG Ser	Gly	TCC Ser 820	GTG Val	GTC Val	CTT Leu	Arg	AAC Asn .825	AGC Ser	ACC Thr	5947
ACC Thr	CTG Leu	GTG Val 1830	ATG Met	CAC His	ATG Met	Lys	GTC Val .835	TAT Tyr	GAC Asp	GAG Glu	Ser	ATC Ile 1840	CAG Gln	CTG Leu	GAC Asp	5995 -
His	AAG Lys 1845	GGC Gly	ACC Thr	AAC Asn	Pro	TGC Cys .850	AGT Ser	GTC Val	AAC Asn	Asn	GGT Gly 1855	GAC Asp	TGC Cys	TCC Ser	CAG Gln	6043
CTC Leu 1860	TGC Cys	CTG _. Leu	CCC Pro	Thr	TCA Ser 1865	GAG Glu	ACG Thr	ACC Thr	Arg	TCC Ser 1870	TGC Cys	ATG Met	TGC Cys	Thr	GCC Ala 1875	6091

FIG. 14A

(Sheet 70 of 91)

													•			_	
GIY	ıyı	AGC Ser	Leu	Arg 1880	Ser	GIY	Gin	GIn	Ala 1885	Cys	Glu	Gly	Val	Gly 1890	Ser	6139	
TTT Phe	CTC Leu	CTG Leu	TAC Tyr 1895	TCT Ser	GTG Val	CAT His	Glu	GGA Gly 1900	ATC Ile	AGG Arg	GGA Gly	Ile	CCC Pro 1905	CTG Leu	GAT Asp	6187	
CCC Pro	Asn	GAC Asp 1910	AAG Lys	TCA Ser	GAT Asp	Ala	CTG Leu 1915	GTC Val	CCA Pro	GTG Val	Ser	GGG Gly 920	ACC Thr	TCG Ser	CTG Leu	6235	
Ala	GTC Val 1925	GGC Gly	ATC Ile	GAC Asp	Phe	CAC His 1930	GCT Ala	GAA Glu	AAT Asn	Asp	ACC Thr 1935	ATC Ile	TAC Tyr	TGG Trp	GTG Val	6283	
GAC Asp 1940	ATG Met	GGC Gly	CTG Leu	Ser	ACG Thr 1945	ATC Ile	AGC Ser	CGG Arg	Ala	AAG Lys 1950	CGG Arg	GAC Asp	CAG Gln	Thr	TGG Trp 1955	6331	
CGT Arg	GAA Glu	GAC Asp	Val	GTG Val 1960	ACC Thr	AAT Asn	GGC Gly	Ile	GGC Gly 1965	CGT Arg	GTG Val	GAG Glu	Gly	ATT Ile 1970	GCA Ala	6379	
GTG Val	GAC Asp	TGG Trp	ATC Ile 1975	GCA Ala	GGC Gly	AAC Asn	Ile	TAC Tyr 1980	TGG Trp	ACA Thr	GAC Asp	Gln	GGC Gly 1985	TTT Phe	GAT Asp	6427	
GTC Val	Ile	GAG Glu 1990	GTC Val	GCC Ala	CGG Arg	Leu	AAT Asn 1995	GGC Gly	TCC Ser	TTC Phe	Arg	TAC Tyr	GTG Val	GTG Val	ATC Ile	6475	
Ser	CAG Gln 2005	GGT Gly	CTA Leu	GAC Asp	Lys	CCC Pro	CGG Arg	GCC Ala	ATC Ile	Thr	GTC Val 2015	CAC His	CCG Pro	GAG Glu	AAA Lys	6523	
GGG Gly 2020	TAC Tyr	TTG Leu	TTC Phe	Trp	ACT Thr 2025	GAG Glu	TGG Tṛp	GGT Gly	Gln	TAT Tyr 2030	CCG Pro	CGT Arg	ATT Ile	Glu	CGG Arg 2035	6571	•
TCT Ser	CGG Arg	CTA Leu	Asp	GGC Gly 2040	ACG Thr	GAG Glu	CGT Arg	Val	GTG Val 2045	CTG Leu	GTC Val	AAC Asn	Val	AGC Ser 2050	ATC Ile	6619	
AGC Ser	TGG Trp	CCC Pro	AAC Asn 2055	GGC Gly	ATC Ile	TCA Ser	Val	GAC Asp 2060	TAC Tyr	CAG Gln	GAT Asp	Gly	AAG Lys 2065	CTG Leu	TAC Tyr	6667	
TGG Trp	Cys	GAT Asp 2070	GCA Ala	CGG Arg	ACA Thr	Asp	AAG Lys 2075	ATT Ile	GAA Glu	CGG Arg	Ile	GAC Asp	CTG Leu	GAG Glu	ACA Thr	6715 -	
GTĀ	GAG Glu 2085	AAC Asn	CGC Arg	GAG Glu	Val	GTT Val 2090	CTG Leu	TCC Ser	AGC Ser	Asn	AAC Asn 2095	ATG Met	GAC Aşp	ATG Met	TTT Phe	67,63	
TCA Ser 2100	GTG Val	TCT Ser	GTG Val	Phe	GAG Glu 2105	GAT Asp	TTC Phe	ATC Ile	Tyr	TGG Trp	AGT Ser	GAC Asp	AGG Arg	Thr	CAT His 2115	6811	

FIG. 14A

(Sheet 71 of 91)

GCC Ala	AAC Asn	Gly GGC	Ser	ATC Ile 2120	AAG Lys	CGC Arg	GGG Gly	Ser	AAA Lys 125	GAC Asp	AAT Asn	GCC Ala	Thr	GAC Asp	TCC Ser	6859
GTG Val	CCC Pro	Leu	CGA Arg 2135	ACC Thr	GGC Gly	ATC Ile	Gly	GTC Val	CAG Gln	CTT Leu	AAA Lys	Asp	ATC Ile 2145	AAA Lys	GTC Val	6907
	Asn					Lys					TGC Cys 2					6955
Gly	GGG Gly 2165	TGC Cys	CAG Gln	CAG Gln	Leu	TGC Cys 2170	CTG Leu	TAC Tyr	CGG Arg	Gly	CGT Arg 2175	GGG Gly	CAG Gln	CGG Arg	GCC Ala	7003
TGC Cys 2180	GCC Ala	TGT Cys	GCC Ala	His	GGG Gly 2185	ATG Met	CTG Leu	GCT Ala	Glu	GAC Asp 2190	GGA Gly	GCA Ala	TCG Ser	Cys	CGC Arg 2195	7051
GAG Glu	TAT Tyr	GCC Ala	Gly	TAC Tyr 2200	CTG Leu	CTC Leu	TAC Tyr	Ser	GAG Glu 2205	CGC Arg	ACC Thr	ATT Ile	Leu	AAG Lys 2210	AGT Ser	7099
		Leu					Asn				CCC Pro	Val				7147
	Asp					Lys					CTG Leu					7195
Arg					Pro					Arg	ATC Ile 2255					7243
				Asn					Asn		GAT Asp			Arg		7291
			Val					Ser		Glu	GGC Gly		Ala		His	.7339
		Trp					Trp		Ser		ACG Thr			Thr		7387
	Arg		Thr			Gln		Arg			Ala		Glu		GAG Glu	7435
Thr		Ile			Ser		Asp				_	Ala			TTG Leu	7483
	Glu					Met					Trp				CAT His 2355	7531

			ATG Met 2					Ser					Leu			7579
		Lys	GAC Asp 2375				Pro .					Ile				7627
	Glu		CTC Leu			Ser					Asp					7675
Cys			GAC Asp		Ser					Ile						7723
			TTC Phe	Gly					Gly					Trp		7771
GAC Asp	TGG Trp	GTG Val	CGG Arg	CGG Arg 2440	GCA Ala	GTG Val	CAG Gln	Arg	GCC Ala 2445	AAC Asn	AAG Lys	CAC His	Val	GGC Gly 2450	AGC Ser	7819
AAC Asn	ATG Met	Lys	CTG Leu 2455	CTG Leu	CGC Arg	GTG Val	Asp	ATC Ile 2460	CCC Pro	CAG Gln	CAG Gln	Pro	ATG Met 2465	GGC Gly	ATC Ile	7867
ATC Ile	Ala	GTG Val 2470	GCC Ala	AAC Asn	GAC Asp	Thr	AAC Asn 2475	AGC Ser	TGT Cys	GAA Glu	Leu	TCT Ser 2480	CCA Pro	TGC Cys	CGA Arg	7915
Ile	AAC Asn 2485	Asn	GGT Gly	GGC Gly	Cys	CAG Gln 2490	GAC Asp	CTG Leu	TGT Cys	Leu	CTC Leu 2495	ACT Thr	CAC His	CAG Gln	GGC Gly	7963
CAT His 2500	Val	AAC Asn	TGC Cys	Ser	TGC Cys 2505	CGA Arg	GGG Gly	GGC Gly	Arg	ATC Ile 2510	CTC Leu	CAG Gln	GAT Asp	Asp	CTC Leu 2515	8011
ACC Thr	TGC	CGA Arg	Ala	GTG Val 2520	Asn	TCC Ser	TCT Ser	Cys	CGA Arg 2525	GCA Ala	CAA Gln	GAT Asp	Glu	TTT Phe 2530	GAG Glu	. 8059
TGT Cys	GCC Ala	TAA : neA :	GGC Gly 2535	Glu	TGC Cys	ATC Ile	Asn	TTC Phe 2540	Ser	CTG Leu	ACC Thr	TGC Cys	GAC Asp 2545	G17	GTC Val	8107
CCC Pro	CAC His	TGC Cys 2550	: Lys	GAC Asp	AAG Lys	TCC Ser	GAT Asp 2555	Glu	AAG Lys	CCA	Ser	TAC Tyr 2560	Cys	AAC Asr	TCC Ser	8155 -
CGC Arg	C CGG g Arg 256	g Cys	AAG Lys	· AAG Lys	ACT	TTC Phe 2570	: Arg	CAG Gln	TGC Cys	AGC Ser	AAT Asn 2575	Gly	, Arg	TG: Cy:	r GTG s Val	8203
TC(Se) 258(c Ası	C ATO	G CTO	G TGC	TGC Cys 2585	s Asr	GGG Gly	G GCC / Ala	GAC Asp	GAC Asp 2590	Cys	GGC Gly	G GAT Y Ası	r GG o Gl	C TCT y Ser 2595	8251

FIG. 14A

8449-134 (Sheet 73 of 91)

GAC	GAG	ATC	CCT	TGC	AAC	AAG	ACA	GCC	TGT	GGT	GTG	GGC	GAG	TTC	CGC	8299
Asp	Glu	Ile	Pro 2	Cys 600	Asn	Lys	Thr		Cys 605	Gly	Val	Gly		Phe 610	Arg	
		Asp	GGG Gly 2615				Gly					Cys				8347
	Asp		GAG Glu			Ser					Cys					8395
Cys			TAC Tyr		Arg					Gly						8443
			ACC Thr	Ser					Pro					Asp		8491
			TGT Cys					Asp					Pro			8539
		Pro	AGA Arg 2695				Asn					Pro				8587
TGC Cys	Ile	CCC Pro 2710	ATG Met	AGC Ser	TGG Trp	Thr	TGT Cys 2715	GAC Asp	AAA Lys	GAG Glu	Asp	GAC Asp 2720	TGT Cys	GAA Glu	CAT His	8635
Gly	GAG Glu 2725	Asp	GAG Glu	ACC Thr	His	TGC Cys 2730	AAC Asn	AAG Lys	TTC Phe	Cys	TCA Ser 2735	GAG Glu	GCC Ala	CAG Gln	TTT Phe	8683
GAG Glu 2740	Cys	CAG Gln	AAC Asn	His	CGC Arg 2745	TGC Cys	ATC Ile	TCC Ser	Lys	CAG Gln 2750	Trp	CTG Leu	TGT Cys	Asp	GGC Gly 2755	8731
AGC Ser	GAT Asp	GAC Asp		GGG Gly 2760	Asp	GGC	TCA Ser	Asp	GAG Glu 2765	GCT Ala	GCT Ala	CAC	Cys	GAA Glu 2770	Gly	8779 ·
AAG Lys	ACG Thr	TGC Cys	GGC Gly 2775	Pro	TCC Ser	TCC	TTC Phe	TCC Ser 2780	Cys	CCT Pro	GGC Gly	ACC Thr	CAC His 2785	Val	TGC Cys	8827
GTC Val	CCC Pro	GAG Glu 2790	a Arg	TGG Trp	CTC Leu	TGT Cys	GAC Asp 2795	Gly	GAC Asp	AAA Lys	GAC Asp	TG1 Cys 2800	Ala	GAT Asp	GGT Gly	8875 -
GC# Ala	A GAG A Asp 2809	Glu	G AGC	ATC Ile	GCA Ala	GCT Ala 2810	Gl3	TGC Cys	TTG Leu	TAC Tyr	AAC Asr 2815	Sea	C ACT	TG:	GAC S Asp	8923
GA(Asi 282	Are	r GAO	TTC Phe	C ATO	TGC Cys 2825	Glr	AA(Ası	C CGC	CAG Gln	TG0 Cys 2830	: Ile	C CCC	C AAC	G CAG S Hi:	TTC s Phe 2835	8971

(Sheet 74 of 91) 8449-134 GTG TGT GAC CAC GAC CGT GAC TGT GCA GAT GGC TCT GAT GAG TCC CCC 9019 Val Cys Asp His Asp Arg Asp Cys Ala Asp Gly Ser Asp Glu Ser Pro 2840 2845 GAG TGT GAG TAC CCG ACC TGC GGC CCC AGT GAG TTC CGC TGT GCC AAT 9067 Glu Cys Glu Tyr Pro Thr Cys Gly Pro Ser Glu Phe Arg Cys Ala Asn 2860 2855 GGG CGC TGT CTG AGC TCC CGC CAG TGG GAG TGT GAT GGC GAG AAT GAC 9115 Gly Arg Cys Leu Ser Ser Arg Gln Trp Glu Cys Asp Gly Glu Asn Asp 2875 TGC CAC GAC CAG AGT GAC GAG GCT CCC AAG AAC CCA CAC TGC ACC AGC 9163 Cys His Asp Gln Ser Asp Glu Ala Pro Lys Asn Pro His Cys Thr Ser 2890 CCA GAG CAC AAG TGC AAT GCC TCG TCA CAG TTC CTG TGC AGC AGT GGG 9211 Pro Glu His Lys Cys Asn Ala Ser Ser Gln Phe Leu Cys Ser Ser Gly 2905 2910 CGC TGT GTG GCT GAG GCA CTG CTC TGC AAC GGC CAG GAT GAC TGT GGC 9259 Arg Cys Val Ala Glu Ala Leu Leu Cys Asn Gly Gln Asp Asp Cys Gly 2920 2925 GAC AGC TCG GAC GAG CGT GGC TGC CAC ATC AAT GAG TGT CTC AGC CGC 9307 Asp Ser Ser Asp Glu Arg Gly Cys His Ile Asn Glu Cys Leu Ser Arg 2935 2940 AAG CTC AGT GGC TGC AGC CAG GAC TGT GAG GAC CTC AAG ATC GGC TTC 9355 Lys Leu Ser Gly Cys Ser Gln Asp Cys Glu Asp Leu Lys Ile Gly Phe 2955 2950 AAG TGC CGC TGT CGC CCT GGC TTC CGG CTG AAG GAT GAC GGC CGG ACG 9403 Lys Cys Arg Cys Arg Pro Gly Phe Arg Leu Lys Asp Asp Gly Arg Thr 2970 2965 9451 . TGT GCT GAT GTG GAC GAG TGC AGC ACC TTC CCC TGC AGC CAG CGC Cys Ala Asp Val Asp Glu Cys Ser Thr Thr Phe Pro Cys Ser Gln Arg 2995 2985 2990 2980 TGC ATC AAC ACC CAT GGC AGC TAT AAG TGT CTG TGT GTG GAG GGC TAT 9499 Cys Ile Asn Thr His Gly Ser Tyr Lys Cys Leu Cys Val Glu Gly Tyr 9547 GCA CCC CGC GGC GGC GAC CCC CAC AGC TGC AAG GCT GTG ACT GAC GAG

Ala	Pro		Gly		Pro	Ser 3020	Cys	Lys	Ala		Thr 3025	Asp	Glu	
GAA Glu	Pro				Ala				Leu					9595 -
Leu				Tyr				Gln		CTG Leu				9643
GTT Val			Phe		Tyr		Gln							- 9691

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•			-										•			
GTG Val	ACC Thr	ACC Thr	Gln	GGC Gly 080	AGC Ser	ATG Met	ATC Ile	Arg	AGG Arg 8085	ATG Met	CAC His	CTT Leu	AAC Asn 3	GGG Gly 090	AGC Ser	9739
		Gln					Thr					Pro	GAT Asp 3105			9787
	Val					Gly					Cys		AAA Lys			9835
Asp					Ser					Ala			ACG Thr			9883
				Leu					Ala				GAT Asp	Val		9931
			Leu					Trp					CTG Leu			9979
		Gly					Ser					Val	GAC Asp 3185			10027
	Thr					Leu					Val		GAG Glu			10075
Tyr					Arg					Glu			AGC Ser			10123
				His					Gln				CAC His	Ile		10171
			Leu							Trp			TGG Trp			10219
		Ile							Thr				AAA Lys 3265	Thr		10267
			Thr					Met					TTC Phe			10315
		Gln					Asn					· Val	AAC Asn		GGT	10363
	Cys					Leu					/ GJ2				TGT Cys 3315	10411

FIG. 14A

8449-134

(Sheet 76 of 91)

•			_										•			
GCC Ala	TGC Cys	CCC Pro	Thr	AAC Asn 320	TTC Phe	TAC Tyr	CTG Leu	Gly :	AGC (Ser 1 325	GAT (GGG Gly	CGC Arg	Thr	TGT Cys 330	GTG Val	10459
		TGC Cys 3					Phe					Asp				10507
CCC Pro	Phe	TGG Trp 3350	TGG Trp	AAG Lys	TGT Cys	Asp	ACC Thr 355	GAG Glu	GAC Asp	GAC Asp	Cys	GGG Gly 360	GAC Asp	CAC His	TCA Ser	10555
Asp	GAG Glu 3365	CCC Pro	CCG Pro	GAC Asp	Cys	CCT Pro 3370	GAG Glu	TTC Phe	AAG Lys	Cys	CGG Arg 375	CCC Pro	GGA Gly	CAG Gln	TTC Phe	10603
CAG Gln 3380	TGC Cys	TCC Ser	ACA Thr	Gly	ATC Ilė 3385	TGC Cys	ACA Thr	AAC Asn	Pro	GCC Ala 390	TTC Phe	ATC Ile	TGC Cys	Asp	GGC Gly 3395	10651
GAC Asp	AAT Asn	GAC Asp	Cys	CAG Gln 3400	GAC Asp	AAC Asn	AGT Ser	Asp	GAG Glu 8405	GCC Ala	AAC Asn	TGT Cys	Asp	ATC Ile 3410	CAC His	10699
GTC Val	TGC Cys	Leu	CCC Pro 3415	AGT Ser	CAG Gln	TTC Phe	Lys	TGC Cys 3420	ACC Thr	AAC Asn	ACC Thr	Asn	CGC Arg 3425	TGT Cys	ATT Ile	10747
CCC Pro	Gly	ATC Ile 3430	Phe	CGC Arg	TGC Cys	Asn	GGG Gly 3435	CAG Gln	GAC Asp	AAC Asn	Cys	GGA Gly 3440	GAT Asp	GGG Gly	GAG Glu	10795
Asp	GAG Glu 3445	Arg	GAC Asp	TGC Cys	Pro	GAG Glu 3450	GTG Val	ACC Thr	TGC Cys	Ala	CCC Pro 3455	AAC Asn	CAG Gln	TTC Phe	CAG Gln	10843
TGC Cys 3460	Ser	ATT	ACC Thr	Lys	CGG Arg 3465	Cys	ATC Ile	CCC Pro	Arg	GTC Val 3470	Trp	GTC Val	TGC Cys	GAC Asp	CGG Arg 3475	10891
GAC Asp	AAT Asn	GAC Asp	TGT Cys	GTG Val 3480	Asp	GGC Gly	AGT Ser	Asp	GAG Glu 3485	Pro	GCC Ala	AAC Asn	TGC Cys	ACC Thi 3490	CAG Gln	10939
ATG Met	ACC Thr	TGI Cys	GG1 Gly 3495	v Val	GAC Asp	GAG Glu	TTC Phe	CGC Arg 3500	Cys	AAG Lys	GAT Asp	TCG Ser	GGC Gly 3505	Arq	TGC G Cys	10987
ATC Ile	CCP Pro	A GC0 Ala 3510	a Arg	TGG Trp	AAC Lys	TGI Cys	GAC S Asp 3515	Gly	GAG Glu	GAT Asp	GAC Asp	TG1 Cys 3520	GI?	G GA' / As	r GGC p Gly	11035
TC(Se	G GA: Asj 352	o Glu	G CCC	C AAC b Lys	G GAJ	A GA(1 Gl) 353(ı Cy:	r GAT s Asp	GAP Glu	CGC Arg	2 ACC 3 Thi 353	Cy:	r GAG	G CC.	A TAC o Tyr	11083
CA(Gl: 354	n Ph	C CGG	C TGG g Cy	C AAG s Ly:	354	n As	c cg n Ar	C TGG g Cy:	C GT(s Val	G CC0 L Pro 3550	o Gl	C CG y Ar	C TG	G CA p Gl	G TGC n Cys 3555	11131

FIG. 14A

8449-134

(Sheet 77 of 91)

GAC Asp	TÁC Tyr	GAC Asp	Asn	GAT Asp 560	TGC Cys	GGT Gly	GAC Asp	Asn	TCC Ser 565	GAT Asp	GAA Glu	GAG Glu	Ser	TGC Cys 570	ACC Thr	11179
CCT Pro	CGG Arg	Pro	TGC Cys 3575	TCC Ser	GAG Glu	AGT Ser	Glu	TTC Phe 580	TCC Ser	TGT Cys	GCC Ala	Asn	GGC Gly 3585	CGC Arg	TGC Cys	11227
	GCG Ala					Cys					Asp					11275
Ser	GAC Asp 8605				Cys					Asp						11323
TGC Cys 3620	AAG Lys	AGC Ser	GGC Gly	His	TGC Cys 8625	ATC Ile	CCC Pro	CTG Leu	Arg	TGG Trp 3630	CGC Arg	TGT Cys	GAC Asp	Ala	GAC Asp 3635	11371
	GAC Asp		Met					Glu					Thr			11419
	ACC Thr	Cys					Phe					Thr				11467
	CTG Leu					Asp					Cys					11515
Asp	GAG Glu 3685				Glu					Val						11563
	TTC Phe			Lys					Cys					Arg	CAA Gln 3715	11611
	GAT Asp		Thr					Asp					Glu		_	11659
	CCC Pro	Pro					Thr		Cys			Lys		Glu		11707
	Cys		Asn			Cys		Ser			Leu		Cys		ATG Met	11755
		Asp					Ser					Cys			GAC Asp	11803
	Lys					Ala					Ile				GAG Glu 3795	11851

(Sheet 78 of 91)

844	9-1	34
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044	י דכי	JT														
GCA Ala	CGC Arg	TGC Cys	Val	CGC Arg 8800	ACC Thr	GAG Glu	AAA Lys	Ala	GCC Ala 805	TAC Tyr	TGT Cys	GCC Ala	Cys	CGC Arg 810	TCG Ser	11899
GGC GGC	TTC Phe	His	ACC Thr 8815	GTG Val	CCC Pro	GGC Gly	Gln	CCC Pro 8820	GGA Gly	TGC Cys	CAA Gln	Asp	ATC Ile 8825	AAC Asn	GAG Glu	11947
TGC Cys	Leu	CGC Arg 8830	TTC Phe	GGC Gly	ACC Thr	Cys	TCC Ser 835	CAG Gln	CTC Leu	TGC Cys	AAC Asn 3	AAC Asn 840	ACC Thr	AAG Lys	GGC Gly	11995
Gly	CAC His 8845	CTC Leu	TGC Cys	AGC Ser	Cys	GCT Ala 8850	CGG Arg	AAC Asn	TTC Phe	Met	AAG Lys 8855	ACG Thr	CAC His	AAC Asn	ACC Thr	12043
TGC Cys 3860	AAG Lys	GCC Ala	GAA Glu	Gly	TCT Ser 8865	GAG Glu	TAC Tyr	CAG Gln	Val	CTG Leu 8870	TAC Tyr	ATC Ile	GCT Ala	Asp	GAC Asp 3875	12091
AAT Asn	GAG Glu	ATC Ile	Arg	AGC Ser 8880	CTG Leu	TTC Phe	CCC Pro	Gly	CAC His 8885	CCC Pro	CAT His	TCG Ser	Ala	TAC Tyr 3890	GAG Glu	12139
CAG Gln	GCA Ala	Phe	CAG Gln 3895	GGT Gly	GAC Asp	GAG Glu	Ser	GTC Val 3900	CGC Arg	ATT Ile	GAT Asp	Ala	ATG Met 3905	GAT Asp	GTC Val	12187
	Val					Val					TGG Trp					12235
Ile					Leu					Pro	CCT Pro 3935					12283
				Gln					Val		CAC His			Ile		12331
			Met					Ala			TGG Trp		Ala			12379
		Trp		Asp			Arg				GAG Glu	Val		Gln		12427
AAG Lys	Gly	GAG Glu 3990	AAC Asn	CGC Arg	AAG Lys	Thr	CTC Leu 3995	Ile	TCG Ser	GGC Gly	Met	ATT Ile 4000	Asp	GAG Glu	CCC Pro	12475
His					Asp					Thr		Tyr			GAC Asp	12523
	Gly			Pro		Ile					Met				CTT Leu 4035	12571

FIG. 14A

8449-134 (Sheet 79 of 91)

CGG Arg	GAG Glu	ACA Thr	Leu	GTG Val 040	CAG Gln	GAC Asp	AAC Asn	Ile	CAG Gln 045	TGG Trp	CCC Pro	ACA Thr	GGC Gly 4	CTG Leu 050	GCC Ala	12619
GTG Val	GAT Asp	Tyr	CAC His 1055	AAT Asn	GAG Glu	CGG Arg	Leu	TAC Tyr 1060	TGG Trp	GCA Ala	GAC Asp	Ala	AAG Lys 1065	CTT Leu	TCA Ser	12667
GTC Val	Ile	GGC Gly 1070	AGC Ser	ATC Ile	CGG Arg	Leu	AAT Asn 1075	GGC Gly	ACG Thr	GAC Asp	Pro	ATT Ile 080	GTG Val	GCT Ala	GCT Ala	12715
Asp	AGC Ser 1085	AAA Lys	CGA Arg	GGC Gly	Leu	AGT Ser 1090	CAC His	CCC Pro	TTC Phe	Ser	ATC Ile 1095	GAC Asp	GTC Val	TTT Phe	GAG Glu	12763
				Gly					Asn				TTC Phe	Lys		12811
			Gly					Val					GGC Gly			12859
		Ser					Tyr					Gln	CCC Pro 4145			12907
	Asn					Lys					Leu		CTG Leu			12955
Pro					Cys					Gly			CTG Leu			13003
				Pro					Thr				GAT Asp	Ala		13051 •
			Thr							Asn			AGC Ser		Phe	13099
		Ala		Arg					Arg				CGC Arg 4225	Tyr	ACG Thr	13147
	Asp		Cys					Cys			His		Arg		GGG	13195
		Cys					Ser					Cys			CCC Pro	13243
	Gly					Lys					val				TAC Tyr 4275	13291

FIG. 14A

•			-													
TGT Cys	GCC Ala	AAC Asn	Asn	AGC Ser 280	ACC Thr	TGC Cys	ACT Thr	Val	AAC Asn 285	CAG Gln	GGC Gly	AAC Asn	Gln	CCC Pro 290	CAG Gln	13339
TGC Cys	CGA Arg	TGC Cys	CTA Leu 295	CCC Pro	GGC Gly	TTC Phe	Leu	GGC Gly 300	GAC Asp	CGC Arg	TGC Cys	Gln	TAC Tyr 1305	CGG Arg	CAG Gln	13387
	Ser	GGC Gly 4310				Asn					Gln					13435
Gly		CGA Arg			Arg					Phe						13483
		AAC Asn		Cys					Glu					Val		13531
		AGT Ser	Gly					Asn					Arg			13579
		TGT Cys					Gly					Gly				13627
	Met	AAC Asn 4390				Met					Cys					13675
Thr		CCC Pro			Glu					Ser						13723
	Ile	GCC Ala		Ile					Leu					Leu	_	13771
		GCC Ala	Gly					Tyr		Arg			Gln		Ala	13819
		TTC Phe		His			Met		Asn					Val		13867
		AAC Asn 4470	Pro			Lys		Tyr			Gly		Pro		GAT Asp	13915
		Gly					Asp					Pro			CCC Pro	139 <u>6</u> 3
	Asr					Val					Tyr				C CAT His 4515	14011

FIG. 14A

8449-134 (Sheet 81 of 91)

GGC AGT CGC CAC TCC CTG GCC AGC ACG GAC GAG AAG CGA GAA CTC CTG

Gly Ser Arg His Ser Leu Ala Ser Thr Asp Glu Lys Arg Glu Leu Leu

4520

4525

4530

GGC CGG GGC CCT GAG GAC GAG ATA GGG GAC CCC TTG GCA TAGGGCCCTG CC 14110 CCGTCGGACT GCCCCAGAA AGCCTCCTGC CCCCTGCCGG TGAAGTCCTT CAGTGAGCCC 14170 Gly Arg Gly Pro Glu Asp Glu Ile Gly Asp Pro Leu Ala 4535 4540

CTCCCCAGCC AGCCCTTCCC	TGGCCCCGCC	GGATGTATAA	ATGTAAAAAT	GAAGGAATTA	14230
CATTTTATAT GTGAGCGAGC	AAGCCGGCAA	GCGAGCACAG	TATTATTTCT	CCATCCCCTC	14290
CCTGCCTGCT CCTTGGCACC	CCCATGCTGC	CTTCAGGGAG	ACAGGCAGGG	AGGGCTTGGG	14350
GCTGCACCTC CTACCCTCCC	ACCAGAACGC	ACCCCACTGG	GAGAGCTGGT	GGTGCAGCCT	14410
TCCCCTCCCT GTATAAGACA	CTTTGCCAAG	GCTCTCCCCT	CTCGCCCCAT	CCCTGCTTGC	14470
CCGCTCCCAC AGCTTCCTGA	GGGCTAATTC	TGGGAAGGGA	GAGTTCTTTG	CTGCCCCTGT	14530
CTGGAAGACG TGGCTCTGGG	TGAGGTAGGC	GGGAAAGGAT	GGAGTGTTTT	AGTTCTTGGG	14590
GGAGGCCACC CCAAACCCCA	GCCCCAACTC	CAGGGGCACC	TATGAGATGG	CCATGCTCAA	14650
CCCCCTCCC AGACAGGCCC	TCCCTGTCTC	CAGGGCCCCC	ACCGAGGTTC	CCAGGGCTGG	14710
AGACTTCCTC TGGTAAACAT	TCCTCCAGCC	TCCCCTCCCC	TGGGGACGCC	AAGGAGGTGG	14770
GCCACACCCA GGAAGGGAAA	GCGGGCAGCC	CCGTTTTGGG	GACGTGAACG	TTTTAATAAT	14830
TTTTGCTGAA TTCTTTACAA	CTAAATAACA	CAGATATTCT	TATAAATAAA	ATTGTAAAAA	14890
AAAAA					14896

Met	Leu	Thr	Pro	Pro	Leu	Leu	Leu	Leu	Leu	Pro	Leu	Leu	Ser	Ala	Leu
T				5					10				Lys	15	
	٠٠: ث		20145	· · · · · · · · · · · · · · · · · · ·				つちょう					20		
Ala,	Cys	Arg 35	Asp	Gln	Ile	Thr,	Cys 40	Ile	Ser.	Lys	Gly.	Trp	Arg	Cys	Asp
Gly	Glu	Arg	Asp	Сув	Pro	Asp	Gly	Ser	Asp	Ğlū	Ala	Pro	Glu	Ile	Cys
Pro	\50 .Gln\	Ser	Lvs	Ala	Gln	Ara	Cvs	Gin	Pro	Aen	.60	Hie	Asn		T ON
65%					,70%	(A				7.5		1120		, S	- 08
GIA	Inr	GIU		Cys 85	yaı	Pro	Met	Ser	Arg 90 a	Leu	Cys	Asn	Gly	yal:	Gln
Asp	Cys	Met	Asp	Gly	Ser.	Asp	Glu	Gly	Pro	His	Cys	Arg	Glu	Leu	Gln
Gly	Asn	Cys	Ser	Arg	Leu	Gly	Cys	105 Gln	His	His	Cvs	Val	110 Pro	Thr	Leu
		115					120					125			
	130					135					140		Gln		
145					150					155			Gly		160
				165					170				Gly	175	
Glu	Gly	Tyr	Leu 180	Leu	Gln	Pro	Asp	Asn 185	Arg	Ser	Cys	Lys	Ala 190	Lys	Asn
Glu	Pro	Val 195	Asp	Arg	Pro	Pro	Val 200	Leu	Leu	Ile	Ala	Asn 205	Ser	Gln	Asn
Ile	Leu 210	Ala	Thr	Tyr	Leu	Ser 215	Gly	Ala	Gln	Val	Ser 220	Thr	Ile	Thr	Pro
Thr 225	Ser	Thr	Arg	Gln	Thr 230	Thr	Ala	Met	Asp	Phe 235	Ser	Tyr	Ala	Asn	Glu 240
				245					250	Ala			Thr	255	Leu
Lys	Cys	Ala	Arg 260	Met	Pro	Gly	Leu	Lys 265	Gly	Phe	Val	Asp	Glu 270	His	Thr
Ile	Asn	Ile 275	Ser	Leu	Ser	Leu	His 280	His	Val	Glu	Gln	Met 285	Ala	Ile	Asp
Trp	Leu 290	Thr	Gly	Asn	Phe	Tyr 295	Phe	Val	Asp	Asp	Ile 300	Asp	Asp	Arg	Ile
Phe 305	Val	Cys	Asn	Arg	Asn 310	Gly	Asp	Thr	Cys	Val 315	Thr	Leu	Leu	Asp	Leu 320
Glu	Leu	Tyr	Asn	Pro 325	Lys	Gly	Ile	Ala	Leu 330	Asp	Pro	Ala	Met	Gly 335	Lys
Val	Phe	Phe	Thr 340	Asp	Tyr	Gly	Gln	Ile 345	Pro	Lys	Val	Glu	Arg 350	Суѕ	Asp
Met	Asp	Gly 355	Gln	Asn	Arg	Thr	Lys 360		Val	Asp	Ser	Lys 365		Val	Phe
Pro	His 370		Ile	Thr	Leu	Asp 375		Val	Ser	Arg	Leu 380		Tyr	Trp	Ala
Asp 385	Ala	Tyr	Leu	Asp	Tyr 390	Ile	Glu	Val	Val	Asp 395		Glu	Gly	Lys	Gly 400
Arg	Gln	Thr	Ile	Ile 405		Gly	Ile	Leu	Ile 410		His	Leu	Tyr	Gly 415	Leu
Thr	Val	Phe	Glu 420		Tyr	Leu	Tyr	Ala 425		Asn	Ser	Asp	Asn 430	Ala	
Ala	Gln	Gln 435		Thr	Ser	Val	Ile 440		Val	Asn	Arg	Phe 445	Asn	Ser	Thr
Glu	Tyr 450		Val	Val	Thr	Arg 455		Asp	Lys	Gly	Gly 460		Leu	His	Ile

FIG. 14B

```
Tyr His Gln Arg Arg Gln Pro Arg Val Arg Ser His Ala Cys Glu Asn
                                           470
                                                                                     475
Asp Gln Tyr Gly Lys Pro Gly Gly Cys Ser Asp Ile Cys Leu Leu Ala
                                   485
                                                                             490
Asn Ser His Lys Ala Arg Thr Cys Arg Cys Arg Ser Gly Phe Ser Leu
                          500
                                                                     505
Gly Ser Asp Gly Lys Ser Cys Lys Lys Pro Glu His Glu Leu Phe Leu
                                                            520
                                                                                                       525
Val Tyr Gly Lys Gly Arg Pro Gly Ile Ile Arg Gly Met Asp Met Gly
                                                    535
                                                                                              540
Ala Lys Val Pro Asp Glu His Met Ile Pro Ile Glu Asn Leu Met Asn
                                           550
                                                                                     555
Pro Arg Ala Leu Asp Phe His Ala Glu Thr Gly Phe Ile Tyr Phe Ala
                                   565
                                                                              570
Asp Thr Thr Ser Tyr Leu Ile Gly Arg Gln Lys Ile Asp Gly Thr Glu
                          580
                                                                     585
Arg Glu Thr Ile Leu Lys Asp Gly Ile His Asn Val Glu Gly Val Ala
                                                            600
                                                                                                       605
Val Asp Trp Met Gly Asp Asn Leu Tyr Trp Thr Asp Asp Gly Pro Lys
                                                    615
                                                                                               620
Lys Thr Ile Ser Val Ala Arg Leu Glu Lys Ala Ala Gln Thr Arg Lys
                                            630
625
Thr Leu Ile Glu Gly Lys Met Thr His Pro Arg Ala Ile Val Val Asp
                                   645
                                                                              650
Pro Leu Asn Gly Trp Met Tyr Trp Thr Asp Trp Glu Glu Asp Pro Lys
                                                                      665
Asp Ser Arg Arg Gly Arg Leu Glu Arg Ala Trp Met Asp Gly Ser His
                                                             680
Arg Asp Ile Phe Val Thr Ser Lys Thr Val Leu Trp Pro Asn Gly Leu
                                                     695
                                                                                                700
Ser Leu Asp Ile Pro Ala Gly Arg Leu Tyr Trp Val Asp Ala Phe Tyr
                                           710
                                                                                      715
Asp Arg Ile Glu Thr Ile Leu Leu Asn Gly Thr Asp Arg Lys Ile Val
                                   725
                                                                              730
 Tyr Glu Gly Pro Glu Leu Asn His Ala Phe Gly Leu Cys His His Gly
                           740
                                                                      745
 Asn Tyr Leu Phe Trp Thr Glu Tyr Arg Ser Gly Ser Val Tyr Arg Leu
                                                             760
                                                                                                         765
 Glu Arg Gly Val Gly Gly Ala Pro Pro Thr Val Thr Leu Leu Arg Ser
                                                     775
                                                                                                780
         770
 Glu Arg Pro Pro Ile Phe Glu Ile Arg Met Tyr Asp Ala Gln Gln
                                            790
                                                                                       795
 Gln Val Gly Thr Asn Lys Cys Arg Val Asn Asn Gly Gly Cys Ser Ser
                                   805
                                                                              810
 Leu Cys Leu Ala Thr Pro Gly Ser Arg Gln Cys Ala Cys Ala Glu Asp
                                                                      825
                                                                                                                  830
 Gln Val Leu Asp Ala Asp Gly Val Thr Cys Leu Ala Asn Pro Ser Tyr
 Val Pro Pro Pro Gln Cys Gln Pro Gly Glu Phe Ala Cys Ala Asn Ser

850 855

Arg Cys Ile Gln Glu Arg Trp Lys Cys Asp Gly Asn Asn Asp Cys Leu

865 870 875
865
| 870 | 875 | 875 | 880 | 885 | 885 | 890 | 895 | 890 | 895 | 895 | 890 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 | 895 |
```

FIG. 14B

| Ala | Thr | Cys | Ser | Ala | Arg | Thr | Cys | Pro | Pro | Asn | Gln | Phe | Ser | Cys | Ala | 930 | 935 | 940 | 940 | 945 | 950 | 955 | 955 | 960 | 955 | 960 | 955 | 960 | 970 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 | 975 Ala Thr Cys Ser Ala Arg Thr Cys Pro Pro Asn Gln Phe Ser Cys Ala 1110 1115 1120 Lys Ala Trp Val Cys Asp Gly Asp Asn Asp Cys Glu Asp Asn Ser Asp 1125 Glu Glu Asn Cys Glu Ser Leu <u>Ala Cys Arq Pro Pro Ser His Pro Cys</u> 1140 1145 1150 Ala Asn Asn Thr Ser Val Cys Leu Pro Pro Asp Lys Leu Cys Asp Gly
1155 1160 1165 Asn Asp Asp Cys Glv Asp Glv Ser Asp Glu Glv Glu Leu Cvs Asp Gln 1175 1180 Cys Ser Leu Asn Asn Gly Gly Cys Ser His Asn Cys Ser Val Ala Pro 1190 1195 Gly Glu Gly Ile Val Cys Ser Cys Pro Leu Gly Met Glu Leu Gly Pro 1205 1210 1215 Asp Asn His Thr Cys Gln Ile Gln Ser Tyr Cys Ala Lys His Leu Lys 1225 1230 1220 Cys Ser Gln Lys Cys Asp Gln Asn Lys Phe Ser Val Lys Cys Ser Cys 1240 1245 Tyr Glu Gly Trp Val Leu Glu Pro Asp Gly Glu Ser Cys Arg Ser Leu 1250 1260 1250 1255 1260 Asp Pro Phe Lys Pro Phe Ile Ile Phe Ser Asn Arg His Glu Ile Arg 1270 1275 Arg Ile Asp Leu His Lys Gly Asp Tyr Ser Val Leu Val Pro Gly Leu 1285 1290 1295 Arg Asn Thr Ile Ala Leu Asp Phe His Leu Ser Gln Ser Ala Leu Tyr 1300 1305 1310 Trp Thr Asp Val Val Glu Asp Lys Ile Tyr Arg Gly Lys Leu Leu Asp 1315 1320 1325 Asn Gly Ala Leu Thr Ser Phe Glu Val Val Ile Gln Tyr Gly Leu Ala 1335 Thr Pro Glu Gly Leu Ala Val Asp Trp Ile Ala Gly Asn Ile Tyr Trp 1355 - 1360 1350 Val Glu Ser Asn Leu Asp Gln Ile Glu Val Ala Lys Leu Asp Gly Thr 1370 1375 1365 Leu Arg Thr Thr Leu Leu Ala Gly Asp Ile Glu His Pro Arg Ala Ile 1385 1380

Ala Leu Asp Pro Arg Asp Gly Ile Leu Phe Trp Thr Asp Trp Asp Ala 1395 1400 1405 Ser Leu Pro Arg Ile Glu Ala Ala Ser Met Ser Gly Ala Gly Arg Arg 1410 1415 1420 Thr Val His Arg Glu Thr Gly Ser Gly Gly Trp Pro Asn Gly Leu Thr 1435 1430 Val Asp Tyr Leu Glu Lys Arg Ile Leu Trp Ile Asp Ala Arg Ser Asp 1445 1450 Ala Ile Tyr Ser Ala Arg Tyr Asp Gly Ser Gly His Met Glu Val Leu 1460 1465 1470 Arg Gly His Glu Phe Leu Ser His Pro Phe Ala Val Thr Leu Tyr Gly 1475 1480 1485 Gly Glu Val Tyr Trp Thr Asp Trp Arg Thr Asn Thr Leu Ala Lys Ala 1490 1495 1500 Asn Lys Trp Thr Gly His Asn Val Thr Val Val Gln Arg Thr Asn Thr 1510 1515 1520 Gln Pro Phe Asp Leu Gln Val Tyr His Pro Ser Arg Gln Pro Met Ala 1525 1530 1535 Pro Asn Pro Cys Glu Ala Asn Gly Gly Gln Gly Pro Cys Ser His Leu 1545 1540 1550 Cys Leu Ile Asn Tyr Asn Arg Thr Val Ser Cys Ala Cys Pro His Leu 1555 1560 1565 Met Lys Leu His Lys Asp Asn Thr Thr Cys Tyr Glu Phe Lys Lys Phe 1570 1575 1580 Leu Leu Tyr Ala Arg Gln Met Glu Ile Arg Gly Val Asp Leu Asp Ala 1590 1595 Pro Tyr Tyr Asn Tyr Ile Ile Ser Phe Thr Val Pro Asp Ile Asp Asn 1605 1610 1615 Val Thr Val Leu Asp Tyr Asp Ala Arg Glu Gln Arg Val Tyr Trp Ser 1620 1625 1630 Asp Val Arg Thr Gln Ala Ile Lys Arg Ala Phe Ile Asn Gly Thr Gly 1635 1640 1645 Val Glu Thr Val Val Ser Ala Asp Leu Pro Asn Ala His Gly Leu Ala 1655 1660 Val Asp Trp Val Ser Arg Asn Leu Phe Trp Thr Ser Tyr Asp Thr Asn 1670 1675 Lys Lys Gln Ile Asn Val Ala Arg Leu Asp Gly Ser Phe Lys Asn Ala 1690 1685 1695 Val Val Gln Gly Leu Glu Gln Pro His Gly Leu Val Val His Pro Leu 1700 1705 1710 Arg Gly Lys Leu Tyr Trp Thr Asp Gly Asp Asn Ile Ser Met Ala Asn 1715 1720 1725 Met Asp Gly Ser Asn Arg Thr Leu Leu Phe Ser Gly Gln Lys Gly Pro 1735 1740 Val Gly Leu Ala Ile Asp Phe Pro Glu Ser Lys Leu Tyr Trp Ile Ser 1750 1755 Ser Gly Asn His Thr Ile Asn Arg Cys Asn Leu Asp Gly Ser Gly Leu 1765 1770 1775 Glu Val Ile Asp Ala Met Arg Ser Gln Leu Gly Lys Ala Thr Ala Leu 1780 1785 Ala Ile Met Gly Asp Lys Leu Trp Trp Ala Asp Gln Val Ser Glu Lys 1800 . 1795 1805 Met Gly Thr Cys Ser Lys Ala Asp Gly Ser Gly Ser Val Val Leu Arg 1810 1815 . 1820 Asn Ser Thr Thr Leu Val Met His Met Lys Val Tyr Asp Glu Ser Ile 1830 1835 Gln Leu Asp His Lys Gly Thr Asn Pro Cys Ser Val Asn Asn Gly Asp 1845 -1850 Cys Ser Gln Leu Cys Leu Pro Thr Ser Glu Thr Thr Arg Ser Cys Met

1860 1865 1870 Cys Thr Ala Gly Tyr Ser Leu Arg Ser Gly Gln Gln Ala Cys Glu Gly 1875 1880 1885 Val Gly Ser Phe Leu Leu Tyr Ser Val His Glu Gly Ile Arg Gly Ile 1895 1900 Pro Leu Asp Pro Asn Asp Lys Ser Asp Ala Leu Val Pro Val Ser Gly 905 1910 1915 1920 Thr Ser Leu Ala Val Gly Ile Asp Phe His Ala Glu Asn Asp Thr Ile 1925 1930 1935 Tyr Trp Val Asp Met Gly Leu Ser Thr Ile Ser Arg Ala Lys Arg Asp 1940 1945 1950 Gln Thr Trp Arg Glu Asp Val Val Thr Asn Gly Ile Gly Arg Val Glu 1955 1960 1965 Gly Ile Ala Val Asp Trp Ile Ala Gly Asn Ile Tyr Trp Thr Asp Gln 1970 1975 1980 Gly Phe Asp Val Ile Glu Val Ala Arg Leu Asn Gly Ser Phe Arg Tyr 1990 1995 Val Val Ile Ser Gln Gly Leu Asp Lys Pro Arg Ala Ile Thr Val His 2005 2010 2015 Pro Glu Lys Gly Tyr Leu Phe Trp Thr Glu Trp Gly Gln Tyr Pro Arg 2020 2025 2030 Ile Glu Arg Ser Arg Leu Asp Gly Thr Glu Arg Val Val Leu Val Asn 2040 2045 2035 Val Ser Ile Ser Trp Pro Asn Gly Ile Ser Val Asp Tyr Gln Asp Gly 2050 2055 2060 Lys Leu Tyr Trp Cys Asp Ala Arg Thr Asp Lys Ile Glu Arg Ile Asp 065 2070 2075 2080 Leu Glu Thr Gly Glu Asn Arg Glu Val Val Leu Ser Ser Asn Asn Met 2085 2090 Asp Met Phe Ser Val Ser Val Phe Glu Asp Phe Ile Tyr Trp Ser Asp 2100 2105 2110 Arg Thr His Ala Asn Gly Ser Ile Lys Arg Gly Ser Lys Asp Asn Ala 2115 2120 2125 Thr Asp Ser Val Pro Leu Arg Thr Gly Ile Gly Val Gln Leu Lys Asp 2130 2135 2140 Ile Lys Val Phe Asn Arg Asp Arg Gln Lys Gly Thr Asn Val Cys Ala 145 2150 2155 2160 Val Ala Asn Gly Gly Cys Gln Gln Leu Cys Leu Tyr Arg Gly Arg Gly 2165 2170 2175 Gln Arg Ala Cys Ala Cys Ala His Gly Met Leu Ala Glu Asp Gly Ala 2185 Ser Cys Arg Glu Tyr Ala Gly Tyr Leu Leu Tyr Ser Glu Arg Thr Ile 2195 2200 2205 Leu Lys Ser Ile His Leu Ser Asp Glu Arg Asn Leu Asn Ala Pro Val 2215 2220 2210 Gln Pro Phe Glu Asp Pro Glu His Met Lys Asn Val Ile Ala Leu Ala 225 2230 2235 Phe Asp Tyr Arg Ala Gly Thr Ser Pro Gly Thr Pro Asn Arg Ile Phe 2245 2250 2255 Phe Ser Asp Ile His Phe Gly Asn Ile Gln Gln Ile Asn Asp Asp Gly 2260 2265 2270 Ser Arg Arg Ile Thr Ile Val Glu Asn Val Gly Ser Val Glu Gly Leu 2280 2285 Ala Tyr His Arg Gly Trp Asp Thr Leu Tyr Trp Thr Ser Tyr Thr Thr 2290 2295 2300 Ser Thr Ile Thr Arg His Thr Val Asp Gln Thr Arg Pro Gly Ala Phe 2310 2315 2320 Glu Arg Glu Thr Val Ile Thr Met Ser Gly Asp Asp His Pro Arg Ala 2330

Phe Val Leu Asp Glu Cys Gln Asn Leu Met Phe Trp Thr Asn Trp Asn 2340 2345 2350 Glu Gln His Pro Ser Ile Met Arg Ala Ala Leu Ser Gly Ala Asn Val 2355 2360 2365 Leu Thr Leu Ile Glu Lys Asp Ile Arg Thr Pro Asn Gly Leu Ala Ile 2375 2380 Asp His Arg Ala Glu Lys Leu Tyr Phe Ser Asp Ala Thr Leu Asp Lys 385 2390 2395 2400 Ile Glu Arg Cys Glu Tyr Asp Gly Ser His Arg Tyr Val Ile Leu Lys 2405 2410 2415 Ser Glu Pro Val His Pro Phe Gly Leu Ala Val Tyr Gly Glu His Ile 2420 2425 2430 Phe Trp Thr Asp Trp Val Arg Arg Ala Val Gln Arg Ala Asn Lys His 2435 2440 24452445 Val Gly Ser Asn Met Lys Leu Leu Arg Val Asp Ile Pro Gln Gln Pro 2450 2455 2460 Met Gly Ile Ile Ala Val Ala Asn Asp Thr Asn Ser Cys Glu Leu Ser 2470 2475 Pro Cys Arg Ile Asn Asn Gly Gly Cys Gln Asp Leu Cys Leu Leu Thr 2485 2490 2495 2490 His Gln Gly His Val Asn Cys Ser Cys Arg Gly Gly Arg Ile Leu Gln 2505 2500 2510 Asp Asp Leu Thr Cys Arg Ala Val Asn Ser Ser Cys Arg Ala Gln Asp 2515 2520 2525 Glu Phe Glu Cys Ala Asn Gly Glu Cys Ile Asn Phe Ser Leu Thr Cys 2530 2535 2540 Asp Gly Val Pro His Cys Lys Asp Lys Ser Asp Glu Lys Pro Ser Tyr 545 2550 2555 2560 Cys Asn Ser Arg Arg Cys Lys Lys Thr Phe Arg Gln Cys Ser Asn Gly 2565 2570 2575 Arg Cys Val Ser Asn Met Leu Trp Cys Asn Gly Ala Asp Asp Cys Gly 2580 2590 Asp Gly Ser Asp Glu Ile Pro Cys Asn Lys Thr Ala Cys Gly Val Gly .2600 2605 Glu Phe Arg Cys Arg Asp Gly Thr Cys Ile Gly Asn Ser Ser Arg Cys 2610 2615 2620 Asn Gln Phe Val Asp Cys Glu Asp Ala Ser Asp Glu Met Asn Cys Ser 625 2630 2635 Ala Thr Asp Cys Ser Ser Tyr Phe Arg Leu Gly Val Lys Gly Val Leu 2645 2650 2655 Phe Gln Pro Cys Glu Arg Thr Ser Leu Cys Tyr Ala Pro Ser Trp Val 2660 2665 2670 Cys Asp Gly Ala Asn Asp Cys Gly Asp Tyr Ser Asp Glu Arg Asp Cys 2675 2680 2685 2675 Pro Gly Val Lys Arg Pro Arg Cys Pro Leu Asn Tyr Phe Ala Cys Pro 2695 2700 Ser Gly Arg Cys Ile Pro Met Ser Trp Thr Cys Asp Lys Glu Asp Asp 705 2710 2715 2720 2710 2715 Cys Glu His Gly Glu Asp Glu Thr His Cys Asn Lys Phe Cys Ser Glu 2725 2730 Ala Gln Phe Glu Cys Gln Asn His Arg Cys Ile Ser Lys Gln Trp Leu 2740 2745 2750 Cys Asp Gly Ser Asp Asp Cys Gly Asp Gly Ser Asp Glu Ala Ala His 2755 2760 2765 . Cys Glu Gly Lys Thr Cys Gly Pro Ser Ser Phe Ser Cys Pro Gly Thr 2775 2780 His Val Cys Val Pro Glu Arg Trp Leu Cys Asp Gly Asp Lys Asp Cys 785 2790 2795 2800 Ala Asp Gly Ala Asp Glu Ser Ile Ala Ala Gly Cys Leu Tyr Asn Ser

2805 2810 Thr Cys Asp Asp Arg Glu Phe Met Cys Gln Asn Arg Gln Cys Ile Pro 2820 2825 2830 Lys His Phe Val Cys Asp His Asp Arg Asp Cys Ala Asp Gly Ser Asp 2835 2840 2845 Glu Ser Pro Glu Cys Glu Tyr Pro Thr Cys Gly Pro Ser Glu Phe Arg 2855 2860 Cys Ala Asn Gly Arg Cys Leu Ser Ser Arg Gln Trp Glu Cys Asp Gly 2870 2875 Glu Asn Asp Cys His Asp Gln Ser Asp Glu Ala Pro Lys Asn Pro His 2890 2885 2895 Cys Thr Ser Pro Glu His Lys Cys Asn Ala Ser Ser Gln Phe Leu Cys 2900 2905 2910 Ser Ser Gly Arg Cys Val Ala Glu Ala Leu Leu Cys Asn Gly Gln Asp 2920 2925 Asp Cys Gly Asp Ser Ser Asp Glu Arg Gly Cys His Ile Asn Glu Cys 2930 2940 Leu Ser Arg Lys Leu Ser Gly Cys Ser Gln Asp Cys Glu Asp Leu Lys 2950 2955 Ile Gly Phe Lys Cys Arg Cys Arg Pro Gly Phe Arg Leu Lys Asp Asp 2965 2970 Gly Arg Thr Cys Ala Asp Val Asp Glu Cys Ser Thr Thr Phe Pro Cys 2980 2985 2990 Ser Gln Arg Cys Ile Asn Thr His Gly Ser Tyr Lys Cys Leu Cys Val 2995 3000 3005 Glu Gly Tyr Ala Pro Arg Gly Gly Asp Pro His Ser Cys Lys Ala Val 3010 3015 3020 Thr Asp Glu Glu Pro Phe Leu Ile Phe Ala Asn Arg Tyr Tyr Leu Arg 025 3030 3035 Lys Leu Asn Leu Asp Gly Ser Asn Tyr Thr Leu Leu Lys Gln Gly Leu 3045 3050 3055 Asn Asn Ala Val Ala Leu Asp Phe Asp Tyr Arg Glu Gln Met Ile Tyr 3060 3065 3070 Trp Thr Asp Val Thr Thr Gln Gly Ser Met Ile Arg Arg Met His Leu 3080 3085 Asn Gly Ser Asn Val Gln Val Leu His Arg Thr Gly Leu Ser Asn Pro 3090 3095 3100 Asp Gly Leu Ala Val Asp Trp Val Gly Gly Asn Leu Tyr Trp Cys Asp 105 3110 3115 3120 Lys Gly Arg Asp Thr Ile Glu Val Ser Lys Leu Asn Gly Ala Tyr Arg 3125 3130 Thr Val Leu Val Ser Ser Gly Leu Arg Glu Pro Arg Ala Leu Val Val 3145 3150 Asp Val Gln Asn Gly Tyr Leu Tyr Trp Thr Asp Trp Gly Asp His Ser 3155 3160 3165 3160 3165 Leu Ile Gly Arg Ile Gly Met Asp Gly Ser Ser Arg Ser Val Ile Val 3170 3175 3180 Asp Thr Lys Ile Thr Trp Pro Asn Gly Leu Thr Leu Asp Tyr Val Thr 3190 3195 Glu Arg Ile Tyr Trp Ala Asp Ala Arg Glu Asp Tyr Ile Glu Phe Ala 3205 3210 3215 Ser Leu Asp Gly Ser Asn Arg His Val Val Leu Ser Gln Asp Ile Pro 3220 3225 3230His Ile Phe Ala Leu Thr Leu Phe Glu Asp Tyr Val Tyr Trp Thr Asp 3235 3240 3245 Trp Glu Thr Lys Ser Ile Asn Arg Ala His Lys Thr Thr Gly Thr Asn 3250 3260 Lys Thr Leu Leu Ile Ser Thr Leu His Arg Pro Met Asp Leu His Val

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8449-134 Phe His Ala bed Ary Gin Pro Asp Val Pro Asn His Pro Cys Lys Val 3285 3290 Asn Asn Gly Gly Cys Ser Asn Leu Cys Leu Leu Ser Pro Gly Gly Gly 3300 3305 3310 His Lys Cys Ala Cys Pro Thr Asn Phe Tyr Leu Gly Ser Asp Gly Arg 3315 3320 3325 Thr Cys Val Ser Asn Cys Thr Ala Ser Gln Phe Val Cys Lys Asn Asp 3335 3340 3330 Lys Cys Ile Pro Phe Trp Trp Lys Cys Asp Thr Glu Asp Asp Cys Gly 345 3350 3355 3360 Asp His Ser Asp Glu Pro Pro Asp Cys Pro Glu Phe Lys Cys Arg Pro 3365 3370 3375 Gly Gln Phe Gln Cys Ser Thr Gly Ile Cys Thr Asn Pro Ala Phe Ile 3380 3385 3390 Cys Asp Gly Asp Asn Asp Cys Gln Asp Asn Ser Asp Glu Ala Asn Cys 3395 3400 3405 Asp Ile His Val Cys Leu Pro Ser Gln Phe Lys Cys Thr Asn Thr Asn 3415 3420 Arg Cys Ile Pro Gly Ile Phe Arg Cys Asn Gly Gln Asp Asn Cys Gly 3430 3435 Asp Gly Glu Asp Glu Arg Asp Cys Pro Glu Val Thr Cys Ala Pro Asn 3455 3445 3450 Gln Phe Gln Cys Ser Ile Thr Lys Arg Cys Ile Pro Arg Val Trp Val 3460 3465 3470 3465 3460 3470 Cys Asp Arg Asp Asn Asp Cys Val Asp Gly Ser Asp Glu Pro Ala Asn 3475 3480 3485 Cys Thr Gln Met Thr Cys Gly Val Asp Glu Phe Arg Cys Lys Asp Ser 3490 3495 3500 Gly Arg Cys Ile Pro Ala Arg Trp Lys Cys Asp Gly Glu Asp Asp Cys 3510 3515 505 Gly Asp Gly Ser Asp Glu Pro Lys Glu Glu Cys Asp Glu Arg Thr Cys 3530 3535 3525 Glu Pro Tyr Gln Phe Arg Cys Lys Asn Asn Arg Cys Val Pro Gly Arg 3540 3545 3550 Trp Gln Cys Asp Tyr Asp Asn Asp Cys Gly Asp Asn Ser Asp Glu Glu 3555 3560 3565 Ser Cys Thr Pro Arg Pro Cys Ser Glu Ser Glu Phe Ser Cys Ala Asn 3570 3575 3580 Gly Arg Cys Ile Ala Gly Arg Trp Lys Cys Asp Gly Asp His Asp Cys 3595 3600 Ala Asp Gly Ser Asp Glu Lys Asp Cys Thr Pro Arg Cys Asp Met Asp 3610 3605 Gln Phe Gln Cys Lys Ser Gly His Cys Ile Pro Leu Arg Trp Arg Cys 3620 3625 3630 Asp Ala Asp Ala Asp Cys Met Asp Gly Ser Asp Glu Glu Ala Cys Gly 3640 3645 Thr Gly Val Arg Thr Cys Pro Leu Asp Glu Phe Gln Cys Asn Asn Thr 3655 . 3660 Leu Cys Lys Pro Leu Ala Trp Lys Cys Asp Gly Glu Asp Asp Cys Gly 3675 3670 Asp Asn Ser Asp Glu Asn Pro Glu Glu Cys Ala Arg Phe Val Cys Pro 3690 3685 3695 Pro Asn Arg Pro Phe Arg Cys Lys Asn Asp Arg Val Cys Leu Trp Ile 3700 3705 3710 Gly Arg Gln Cys Asp Gly Thr Asp Asn Cys Gly Asp Gly Thr Asp Glu 3720 3725 3715 Glu Asp Cys Glu Pro Pro Thr Ala His Thr Thr His Cys Lys Asp Lys 3730 3735 3740 Lys Glu Phe Leu Cys Arg Asn Gln Arg Cys Leu Ser Ser Ser Leu Arg

FIG. 14B

745 3750 Cys Asn Met Phe Asp Asp Cys Gly Asp Gly Ser Asp Glu Glu Asp Cys 3765 3770 Ser Ile Asp Pro Lys Leu Thr Ser Cys Ala Thr Asn Ala Ser Ile Cys 3785 Gly Asp Glu Ala Arg Cys Val Arg Thr Glu Lys Ala Ala Tyr Cys Ala 3800 3805 Cys Arg Ser Gly Phe His Thr Val Pro Gly Gln Pro Gly Cys Gln Asp 3815 3820 Ile Asn Glu Cys Leu Arg Phe Gly Thr Cys Ser Gln Leu Cys Asn Asn 3830 3835 Thr Lys Gly Gly His Leu Cys Ser Cys Ala Arg Asn Phe Met Lys Thr 3845 3850 3855 His Asn Thr Cys Lys Ala Glu Gly Ser Glu Tyr Gln Val Leu Tyr Ile 3860 3865 Ala Asp Asp Asn Glu Ile Arg Ser Leu Phe Pro Gly His Pro His Ser 3880 Ala Tyr Glu Gln Ala Phe Gln Gly Asp Glu Ser Val Arg Ile Asp Ala 3895 Met Asp Val His Val Lys Ala Gly Arg Val Tyr Trp Thr Asn Trp His 3900 3910 3915 Thr Gly Thr Ile Ser Tyr Arg Ser Leu Pro Pro Ala Ala Pro Pro Thr 3925 3930 Thr Ser Asn Arg His Arg Arg Gln Ile Asp Arg Gly Val Thr His Leu 3940 3945 Asn Ile Ser Gly Leu Lys Met Pro Arg Gly Ile Ala Ile Asp Trp Val 3960 3965 Ala Gly Asn Val Tyr Trp Thr Asp Ser Gly Arg Asp Val Ile Glu Val 3970 3975 3980 Ala Gln Met Lys Gly Glu Asn Arg Lys Thr Leu Ile Ser Gly Met Ile 3990 3995 Asp Glu Pro His Ala Ile Val Val Asp Pro Leu Arg Gly Thr Met Tyr 4005 4010 Trp Ser Asp Trp Gly Asn His Pro Lys Ile Glu Thr Ala Ala Met Asp 4025 Gly Thr Leu Arg Glu Thr Leu Val Gln Asp Asn Ile Gln Trp Pro Thr 4040 Gly Leu Ala Val Asp Tyr His Asn Glu Arg Leu Tyr Trp Ala Asp Ala 4045 4055 4060 Lys Leu Ser Val Ile Gly Ser Ile Arg Leu Asn Gly Thr Asp Pro Ile 4070 4075 4080 Val Ala Ala Asp Ser Lys Arg Gly Leu Ser His Pro Phe Ser Ile Asp 4085 4090 4095 Val Phe Glu Asp Tyr Ile Tyr Gly Val Thr Tyr Ile Asn Asn Arg Val 4100 4105 Phe Lys Ile His Lys Phe Gly His Ser Pro Leu Val Asn Leu Thr Gly 4115 4120 Gly Leu Ser His Ala Ser Asp Val Val Leu Tyr His Gln His Lys Gln 4125 4135 Pro Glu Val Thr Asn Pro Cys Asp Arg Lys Lys Cys Glu Trp Leu Cys 4150 4155 Leu Leu Ser Pro Ser Gly Pro Val Cys Thr Cys Pro Asn Gly Lys Arg 4165 4170 Leu Asp Asn Gly Thr Cys Val Pro Val Pro Ser Pro Thr Pro Pro Pro 4180 4185 Asp Ala Pro Arg Pro Gly Thr Cys Asn Leu Gln Cys Phe Asn Gly Gly 4190 4200 Ser Cys Phe Leu Asn Ala Arg Arg Gln Pro Lys Cys Arg Cys Gln Pro 4205 4215 4220

Arg Tyr Thr Gly Asp Lys Cys Glu Leu Asp Gln Cys Trp Glu His Cys 4230 4235 Arg Asn Gly Gly Thr Cys Ala Ala Ser Pro Ser Gly Met Pro Thr Cys 4245 4250 Arg Cys Pro Thr Gly Phe Thr Gly Pro Lys Cys Thr Gln Gln Val Cys 4265 4260 4270 Ala Gly Tyr Cys Ala Asn Asn Ser Thr Cys Thr Val Asn Gln Gly Asn 4275 4280 4285 Gln Pro Gln Cys Arg Cys Leu Pro Gly Phe Leu Gly Asp Arg Cys Gln 4295 4300 Tyr Arg Gln Cys Ser Gly Tyr Cys Glu Asn Phe Gly Thr Cys Gln Met 4310 4315 Ala Ala Asp Gly Ser Arg Gln Cys Arg Cys Thr Ala Tyr Phe Glu Gly 4325 4330 4335 Ser Arg Cys Glu Val Asn Lys Cys Ser Arg Cys Leu Glu Gly Ala Cys 4340 4345 4350 Val Val Asn Lys Gln Ser Gly Asp Val Thr Cys Asn Cys Thr Asp Gly 4360 4365 4355 Arg Val Ala Pro Ser Cys Leu Thr Cys Val Gly His Cys Ser Asn Gly 4375 4380 Gly Ser Cys Thr Met Asn Ser Lys Met Met Pro Glu Cys Gln Cys Pro 4390 4395 Pro His Met Thr Gly Pro Arg Cys Glu Glu His Val Phe Ser Gln Gln 4410 4415 4405 Gln Pro Gly His Ile Ala Ser Ile Leu Ile Pro Leu Leu Leu Leu Leu 4420 4425 4430 Leu Leu Val Leu Val Ala Gly Val Val Phe Trp Tyr Lys Arg Arg Val 4435 4440 Gln Gly Ala Lys Gly Phe Gln His Gln Arg Met Thr Asn Gly Ala Met 4455 4460 Asn Val Glu Ile Gly Asn Pro Thr Tyr Lys Met Tyr Glu Gly Gly 4470 4475 Pro Asp Asp Val Gly Gly Leu Leu Asp Ala Asp Phe Ala Leu Asp Pro 4490 4495 4485 Asp Lys Pro Thr Asn Phe Thr Asn Pro Val Tyr Ala Thr Leu Tyr Met 4500 4505 4510 Gly Gly His Gly Ser Arg His Ser Leu Ala Ser Thr Asp Glu Lys Arg 4520 4525 Glu Leu Leu Gly Arg Gly Pro Glu Asp Glu Ile Gly Asp Pro Leu Ala · 4540 4535